The portrait of anxiety

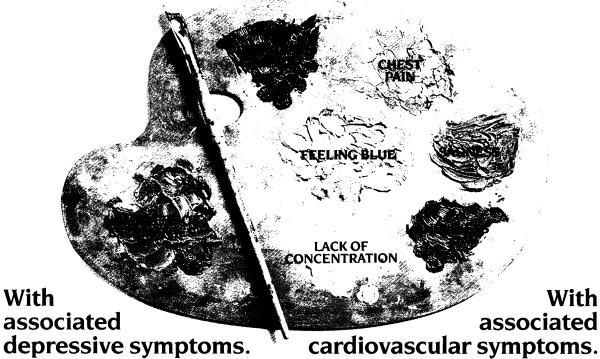


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is often complicated



In double-blind four-week clinical trials in 632 patients with moderate to severe anxiety, therapy with XANAX was compared with placebo.

XANAX was significantly more effective (P<001) than placebo in relieving the anxiety. with over half of the patients showing marked to moderate improvement by the first evaluation period (one week).

In addition, over 70% of these patients

experienced associated moderate to severe depressed mood. XANAX was shown to be significantly more effective (P<.014) than placebo in improving the associated depressed mood.



Almost 60% of patients in the study had anxiety with associated cardiovascular symptoms even though cardiovascular disease had been ruled out. XANAX was shown to effectively relieve anxiety including the associated cardiovascular symptoms.

XANAX the first of a unique class—the

triazolobenzodiazepines.

■ Well tolerated—Side effects. if they occur are generally observed at the beginning of therapy and usually disappear with continued medication. Drowsiness and light-headedness were the most commonly reported adverse reactions.

■ Sustained efficacy—No reported increase in dosage during 16-week clinical study once an appropriate dosage was achieved. Since long-term effectiveness of XANAX has not been established it is recommended that it not be used for longer than 16 weeks.

■ Simple dosage—0.25 to 0.5 mg t.i.d.



for the relief of complicated anxiety

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Hyatt Reger

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Health Management Publications, Inc. will sponsor this unique conference in an effort to create awareness, educate health care professionals, and make a difference in contemporary wound management.

As you'll see, we have assembled the top wound care experts in the country to address the most important topics, procedures, and problems facing today's professionals. Exploring all aspects of pressure ulcers, tissue destruction, the healing process, and the practical aspects of wound pathology/physiology will be of

From these and several other presentations, including the legal and ethical issues surrounding wound management, a protocol for prevention and treatment of pressure ulcers will be developed. You can be part of this history-making group of health care professionals, assembling as a national base to effectively and exclusively examine all problems concerning today's wound care practitioners.

CME INFORMATION:

Extended Programs in Medical Education of the University of California School of Medicine at San Francisco designates this continuing medical education activity for 13 credit hours in Category 1 of the Physician's Recognition Award of the American Medical Association and the Certificate Program of the California Medical Association.

OBJECTIVES:

- Describe current research relative to the wound healing process.
- Identify future trends in wound management.
- Describe the role of biosynthetic coverings in wound healing.
- Discuss the principles of using topical antiseptics for wound management.
- Discuss the legal and ethical issues associated with treating pressure sores.
- Discuss reimbursement/Medicare issues related to the pressure sore patient.
- Identify the essential components of a wound management protocol.
- Discuss current practice in the management of patients with chronic wounds.
- Differentiate between arterial and venous ulcers.
- Identify four factors when choosing the appropriate patient support surface.

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*CAPOTEN® (captopril tablets) may be used as initial therapy only for patients with normal renal function in whom the raisk of neutropenia/agranulocytosis is relatively low (1 out of over 8,600 in clinical trials). Use special precautions white cells or immune response. Evaluation of hypertensives, or those exposed to other drugs known to affect both effects are generally occurring adverse reactions associated with CAPOTEN are skin rash and taste alteration; REACTIONS in the brief summary on the adjacent page.

^{1.} Croog SH, Levine S, Testa MA, et al: The effects of antihypertensive therapy on the quality of life. N Engl J Med





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We spend so much of our lives at work...it's no wonder our work performance is key to our quality of life. Work performance is also a key factor in assessing antihypertensive therapy. CAPOTEN improved hypertensive patients' work performance (e.g., ability to keep pace with the job, concentration, job satisfaction, less on-the-job fatigue). So, for hypertensive patients who work, why not prescribe the antihypertensive that can work for them... CAPOTEN.

These data are based on a multicenter, randomized, 24-week study of 626 mild-to-moderate hypertensive male patients with normal renal function, 181 of whom received captopril.

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CAPOTEN® TABLETS

Captopril Tablets

INDICATIONS: Hypertension—CAPOTEN (captopril) is indicated for the treatment of hypertension. Consideration should be given to the risk of neutropenia/agranulocytosis (see WARNINGS). CAPOTEN may be used as initial therapy for patients with normal renal function, in whom the risk is relatively low. In patients with impaired renal function, particularly those with collagen vascular disease, captopril should be reserved for those who have either developed unacceptable side effects on other drugs, or have failed to respond satisfactorily to drug combinations. CAPOTEN is effective alone and in combination with other antihypertensive agents, especially thiazide-type diuretics.

Heart Failure: CAPOTEN (captopril) is indicated in patients with heart failure who have not responded adequately to or cannot be controlled by conventional diuretic and digitalis therapy. CAPOTEN is to be used with diuretics and digitalis.

CONTRAINDICATIONS: CAPOTEN is contraindicated in patients who are hypersensitive to this product.

WARNINGS: Neutropenia/Agranulocytosis — Neutropenia (< 1000/mm³) with myeloid hypoplasia has resulted from use of captopril. About half of the neutropenic patients developed systemic or oral cavity infections or other features of the syndrome of agranulocytosis. The risk of neutropenia is dependent on the clinical status of the patient:

granulocytosis. The risk of neutropenia is dependent on the clinical status of the patient: In clinical trials in patients with hypertension who have normal renal function (serum creatinine less than 1.6 mg/dL and no collagen vascular disease), neutropenia has been seen in one patient out of over 8,600 exposed. In patients with some degree of renal failure (serum creatinine at least 1.6 mg/dL) but no collagen vascular disease, the risk in clinical trials was about 1 per 500. Doses were relatively high in these patients, particularly in view of their diminished renal function. In patients with collagen vascular diseases (e.g., systemic lupus erythematosus, scleroderma) and impaired renal function, neutropenia occurred in 3.7% of patients in clinical trials. While none of the over 750 patients in formal clinical trials of heart failure developed neutropenia, it has occurred during the subsequent clinical experience. Of reported cases, about half had serum creatinine ≥ 1.6 mg/dL and more than 75% received procainamide. In heart failure, it appears that the same risk factors for neutropenia are present. Seutropenia has appeared usually within 3 months after starting therapy, associated

serum creatinine ≥ 1.0 mg/dL and more than 1/5% received procainamide. In heart failure, it appears that the same risk factors for neutropenia are present.

Neutropenia has appeared usually within 3 months after starting therapy, associated with myeloid hypoplasia and frequently accompanied by erythroid hypoplasia and decreased numbers of megakaryocytes (e.g., hypoplastic bone marrow and pancytopenia); anemia and thrombocytopenia were sometimes seen. Neutrophils generally returned to normal in about 2 weeks after captopril was discontinued, and serious infections were limited to clinically complex patients. About 13% of the cases of neutropenia have ended fatally, but almost all fatalities were in patients with serious illness, having collagen vascular disease, renal failure, heart failure or immunosuppressant therapy, or a combination of these complicating factors. Evaluation of the hypertensive or heart failure patient should always include assessment of renal function. If captopril is used in patients with impaired renal function, white blood cell and differential counts should be evaluated prior to starting treatment and at approximately 2-week intervals for about 3 months, then periodically. In patients with collagen vascular disease or who are exposed to other drugs known to affect the white cells or immune response, particularly when there is impaired renal function, captopril should be used only after an assessment of benefit and risk, and then with caution. All patients treated with captopril should be told to report any signs of infection (e.g., sore throat, fever). If infection is suspected, perform white cell counts without delay. Since discontinuation of captopril and other drugs has generally led to prompt return of the white count to normal, upon confirmation of neutropenia (neutrophil count ≈ 1000/mm³) withdraw captopril and closely follow the patient's course.

Proteinuria: Total urinary proteins ≈ 1g per day were seen in about 0.7% of patients on

Proteinuria: Total urinary proteins ~1 g per day were seen in about 0.7% of patients on captopril. About 90% of affected patients had evidence of prior renal disease or received high doses (>150 mg/day), or both. The nephrotic syndrome occurred in about one-fifth of proteinuric patients. In most cases, proteinuria subsided or cleared within 6 months whether or not captopril was continued. The BUN and creatinine were seldom altered in proteinuric patients. Since most cases of proteinuria occurred by the 8th month of therapy with captopril, patients with prior renal disease or those receiving captopril at doses ~150 mg per day, should have urinary protein estimates (dip-stick on 1st morning urine) before therapy, and periodically thereafter.

Ist morning urine) before therapy, and periodically thereafter.

Hypotension: Excessive hypotension was rarely seen in hypertensive patients but is a possibility in severely salt/volume-depleted persons such as those treated vigorously with diuretics (see PRECAUTIONS [Drug Interactions]). In heart failure, where the blood pressure was either normal or low, transient decreases in mean blood pressure ~20% were recorded in about half of the patients. This transient hypotension may occur after any of the first several doses and is usually well tolerated, although rarely it has been associated with arrhythmia or conduction defects. A starting dose of 6.25 or 12.5 mg it d may minimize the hypotensive effect. Patients should be followed closely for the first 2 weeks of treatment and whenever the dose of captopril and/or diuretic is increased.

BECAUSE OF THE POTENTIAL FALL IN BLOOD PRESSURE IN THESE PATIENTS, THERAPY SHOULD BE STARTED UNDER VERY CLOSE MEDICAL SUPERVISION.

MEDICAL SUPERVISION.

PRECAUTIONS: General: Impaired Renal Function — Hypertension — Some hypertensive patients with renal disease, particularly those with severe renal artery stenosis, have developed increases in BUN and serum creatinine. It may be necessary to reduce captopril dosage and/or discontinue diuretic. For some of these patients, normalization of blood pressure and maintenance of adequate renal perfusion may not be possible. Heart Failure — About 20% of patients develop stable elevations of BUN and serum creatinine > 20% above normal or baseline upon long-term treatment. Less than 5% of patients, generally with severe preexisting renal disease, required discontinuation due to progressively increasing creatinine. See DOSAGE AND ADMINISTRATION, ADVERSE REACTIONS [Altered Laboratory Findings]. Valvular Stenosis — A theoretical concern, for risk of decreased coronary perfusion, has been noted regarding vasodilator treatment in patients with aortic stenosis due to decreased afterload reduction. Surgery Anesthesia — If hypotension occurs during surgery or anesthesia, and is considered due to the effects of captopril, it is correctable by volume expansion.

Drug Interactions: Hypotension — Patients on Diuretic Therapy — Precipitous reduction

Drug Interactions: Hypotension – Patients on Diuretic Therapy – Precipitous reduction of blood pressure may occasionally occur within the 1st hour after administration of the initial of captopril dose in patients on diuretics, especially those recently placed on diuretics, and those on severe dietary salt restriction or dialysis. This possibility can be minimized

by either discontinuing the diuretic or increasing the salt intake about 1 week prior to initiation of captopril therapy or by initiating therapy with small doses (6.25 or 12.5 mg). Alternatively, provide medical supervision for at least 1 hour after the initial dose.

Agents Having Vasodilator Activity – In heart failure patients, vasodilators should be administered with caution.

Agents Causing Renin Release — Captopril's effect will be augmented by antihypertensive agents that cause renin release.

Agents Affecting Sympathetic Activity — The sympathetic nervous system may be especially important in supporting blood pressure in patients receiving captopril alone or with diuretics. Beta-adrenergic blooking drugs add some further antihypertensive effect to captopril, but the overall response is less than additive. Therefore, use agents affecting sympathetic activity (e.g., ganglionic blocking agents or adrenergic neuron blocking agents) with caution.

Agents Increasing Serum Potassium—Give potassium-sparing diuretics or potassium supplements only for documented hypokalemia, and then with caution, since they may lead to a significant increase of serum potassium. Use potassium-containing salt substitutes with caution.

Inhibitors of Endogenous Prostaglandin Synthesis — Indomethacin and other nonsteroidal anti-inflammatory agents may reduce the antihypertensive effect of captopril, especially in low renin hypertension.

Drug/Laboratory Test Interaction: Captopril may cause a false-positive urine test for acetone.

Carcinogenesis, Mutagenesis and Impairment of Fertility: Two-year studies with doses of 50 to 1350 mg/kg/day in mice and rats failed to show any evidence of carcinogenic potential. Studies in rats have revealed no impairment of fertility.

Pregnancy: Category C: There are no adequate and well-controlled studies in pregnant women. Embryocidal effects and craniofacial malformations were observed in rabbits. Therefore, captopril should be used during pregnancy, or for patients likely to become pregnant, only if the potential benefit outweighs the potential risk to the fetus. Captopril crosses the human placenta.

Nursing Mothers: Captopril is secreted in human milk. Exercise caution when administering captopril to a nursing woman, and, in general, nursing should be interrupted.

Pediatric Use: Safety and effectiveness in children have not been established although there is limited experience with use of captopril in children from 2 months to 15 years of age. Dosage, on a weight basis, was comparable to that used in adults. CAPOTEN (captopril) should be used in children only if other measures for controlling blood pressure have not been effective.

ADVERSE REACTIONS: Reported incidences are based on clinical trials involving approximately 7000 patients.

Renal-About 1 of 100 patients developed proteinuria (see WARNINGS). Renal insufficiency, renal failure, polyuria, oliguria, and urinary frequency in 1 to 2 of 1000 patients.

Hematologic — Neutropenia/agranulocytosis has occurred (see WARNINGS). Anemia, thrombocytopenia, and pancytopenia have been reported.

mia, thrombocytopenia, and pancytopenia have been reported.

Dermatologic — Rash, (usually maculopapular, rarely urticarial), often with pruritus, and sometimes with fever and eosinophilia, in about 4 to 7 of 100 patients (depending on renal status and dose), usually during the 1st 4 weeks of therapy. Pruritus, without rash, in about 2 of 100 patients. A reversible associated pemphigoid-like lesion, and photosensitivity, have also been reported. Angioedema of the face, mucous membranes of the mouth, or of the extremities in about 1 of 1000 patients — reversible on discontinguance of captopril therapy. One case of laryngeal edema has been reported. Flushing or pallor in 2 to 5 of 1000 patients.

Cardiovascular — Hypotension may occur; see WARNINGS and PRECAUTIONS [Drug Interactions] for discussion of hypotension on initiation of captopril therapy. Tachycardia, chest pain, and palpitations each in about 1 of 100 patients. Angina pectoris, myocardial infarction, Raynaud's syndrome, and congestive heart failure each in 2 to 3 of 1000 patients.

Dysgusia — Approximately 2 to 4 (depending on renal status and dose) of 100 patients developed a diminution or loss of taste perception; taste impairment is reversible and usually self-limited even with continued drug use (2 to 3 months). Gastric irritation, abdominal pain, nausea, vomiting, diarrhea, anorexia, constipation, aphthous ulcers, peptic ulcer, dizziness, headache, malaise, fatigue, insomnia, dry mouth, dyspnea, cough, alopecia, paresthesias reported in about 0.5 to 2% of patients but did not appear at increased frequency compared to placebo or other treatments used in controlled trials.

Altered Laboratory Findings: Elevations of liver enzymes in a few patients although no causal relationship has been established. Rarely cholestatic jaundice, and hepatocellular injury with or without secondary cholestasis, have been reported. A transient elevation of BUN and serum creatinine may occur, especially in volume-depleted or renovascular hypertension patients. In instances of rapid reduction of longstanding or severely elevated blood pressure, the glomerular filtration rate may decrease transiently, also resulting in transient rises in serum creatinine and BUN. Small increases in serum potassium concentration frequently occur, especially in patients with renal impairment (see PRECAUTIONS).

OVERDOSAGE: Primary concern is correction of hypotension. Volume expansion with an I.V. infusion of normal saline is the treatment of choice for restoration of blood pressure. Captopril may be removed from the general circulation by hemodialysis.

DOSAGE AND ADMINISTRATION: CAPOTEN (captopril) should be taken one hour before meals. In hypertension, CAPOTEN may be dosed bid or tid. Dosage must be individualized; see DOSAGE AND ADMINISTRATION section of package insert for detailed information regarding dosage in hypertension and in heart failure. Because CAPOTEN (captopril) is excreted primarily by the kidneys, dosage adjustments are recommended for patients with impaired renal function.

Consult package insert before prescribing CAPOTEN (captopril).

HOW SUPPLIED: Available in tablets of 12.5, 25, 50, and 100 mg in bottles of 100 (25 mg and 50 mg also available in bottles of 1000), and in UNIMATIC "unit-dose packs of 100 tablets. (J3-658J)



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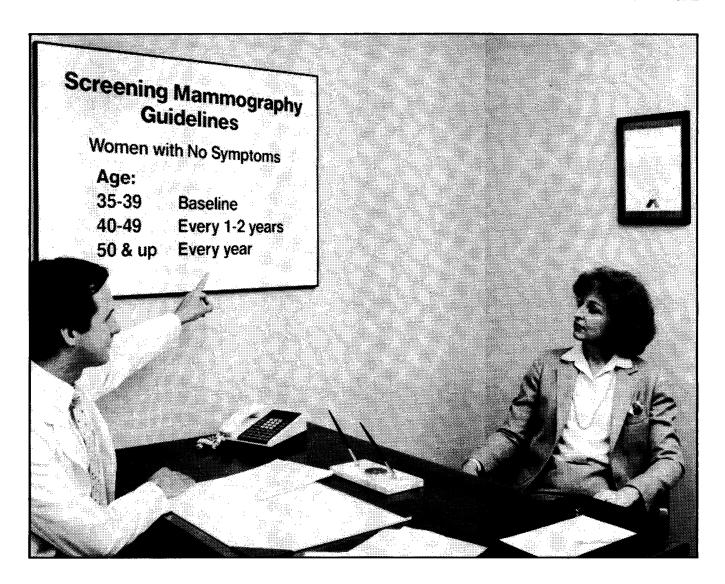


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*CARDIZEM® (dilliazem HCl) is indicated in the treatment of angina pectoris due to coronary artery spasm and in the management of chronic stable angina (classic effort-associated angina) in patients who cannot tolerate therapy with beta-blockers and/or nitrates or who remain symptomatic despite adequate doses of these agents.

[†]See Warnings and Precautions.

Please see brief summary of prescribing information on the next page.

0453S7





ANTIANGINAL PROTECTION PLUS SAFETY diltiazem HCI/Marion

Usual maintenance dosage range: 180-360 mg/day

Brief SummaryProfessional Use Information

CARDIZEM®

(dilfiazem HCl) 30 mg, 60 mg, 90 mg, and 120 mg Tablets

CONTRAINDICATIONS

CARDIZEM is contraindicated in (1) patients with sick sinus syndrome except in the presence of a functioning ventricular pacemaker, (2) patients with second- or third-degree AV block except in the presence of a functioning ventricular pacemaker, and (3) patients with hypotension (less than 90 mm Hg systolic).

WARNINGS

- Cardiac Conduction. CARDIZEM prolongs AV node refractory periods without significantly prolonging sinus node recovery time, except in patients with sick sinus syndrome. This effect may rarely result in abnormally slow heart rates (particularly in patients with sick sinus syndrome) or second- or third-degree AV block (six of 1,243 patients for 0.48%). Concomi-tant use of diltiazem with beta-blockers or digitalis may result in additive effects on cardiac conduction. A patient with Prinzmetal's angina developed periods of asystole (2 to 5 seconds) after a single dose of 60 mg of diltiazem
- **Congestive Heart Failure.** Although diltiazem has a negative inotropic effect in isolated animal tissue preparations, hemodynamic studies in humans with normal ventricular function have not shown a reduction in cardiac index nor consistent negative effects on contractility (dp/dt). Experience with the use of CARDIZEM alone or in combination with beta-blockers in patients with impaired ventricular function is very limited. Caution should be exercised when using the drug in such
- **Hypotension.** Decreases in blood pressure associated with CARDIZEM therapy may occasionally result in symptomatic hypotension.
- Acute Hepatic Injury. In rare instances, significant elevations in enzymes such as alkaline phosphatase, CPK, LDH, SGOT, SGPT, and other symptoms consistent with acute hepatic injury have been noted. These reactions have been reversible upon discontinuation of drug therapy. The relationship to CARDIZEM is uncertain in most cases, but probable in some. (See PRECAUTIONS.)

PRECAUTIONS

General. CARDIZEM (diltiazem hydrochloride) is extensively metabolized by the liver and excreted by the kidneys and in bile. As with any new drug given over prolonged periods, laboratory parameters should be moni-tored at regular intervals. The drug should be used with caution in patients with impaired renal or hepatic function. In subacute and chronic dog and rat studies designed to produce toxicity, high doses of diltiazem were associated with hepatic damage. In special subacute hepatic studies,

oral doses of 125 mg/kg and higher in rats were associated with histological changes in the liver which were reversible when the drug was discontinued. In dogs, doses of 20 mg/kg were also associated with hepatic changes; however, these changes were reversible with continued dosing.

Drug Interaction. Pharmacologic studies indicate that there may be additive effects in prolonging AV conduction when using beta-blockers or digitalis concomitantly with CARDIZEM. (See WARNINGS.) Controlled and uncontrolled domestic studies suggest that

concomitant use of CARDIZEM and beta-blockers or digitalis is usually well tolerated. Available data are not sufficient, however, to predict the effects of concomitant treatment. particularly in patients with left ventricular dysfunction or cardiac conduction abnormalities. In healthy volunteers, diltiazem has been shown to increase serum digoxin levels

Carcinogenesis, Mutagenesis, Impairment of Fertility. A 24-month study in rats and a 21-month study in mice showed no evidence of carcinogenicity. There was also no mutagenic response in in vitro bacterial tests. No intrinsic effect on fertility was observed in rats.

Pregnancy. Category C. Reproduction studies have been conducted in mice, rats, and rabbits. Administration of doses ranging from five to ten times greater (on a mg/kg basis) than the daily recommended therapeutic dose has resulted in embryo and fetal lethality. These doses, in some studies, have been reported to cause skeletal abnormalities. In the perinatal/postnatal studies, there was some reduction in early individual pup weights and survival rates. There was an increased incidence of stillbirths at doses of 20 times the human dose or greater

There are no well-controlled studies in pregnant women, therefore, use CARDIZEM in pregnant women only if the potential benefit justifies the potential risk to the fetus. **Nursing Mothers.** Diltiazem is excreted in human milk.

One report suggests that concentrations in breast milk may approximate serum levels. If use of CARDIZEM is deemed essential, an alternative method of infant feeding should be

Pediatric Use. Safety and effectiveness in children have

ADVERSE REACTIONS

Serious adverse reactions have been rare in studies carried out to date, but it should be recognized that patients with impaired ventricular function and cardiac conduction abnormalities have usually been excluded.

In domestic placebo-controlled trials, the incidence of adverse reactions reported during CARDIZEM therapy was not greater than that reported during placebo therapy.

The following represent occurrences observed in clinical studies which can be at least reasonably associated with the pharmacology of calcium influx inhibition. In many cases, the relationship to CARDIZEM has not been established. The most common occurrences as well as their frequency of presentation are: edema (2.4%), headache (2.1%), nausea (1.9%), dizziness (1.5%), rash (1.3%), asthenia (1.2%). In addition, the following events were reported infrequently (less than 1%):

Cardizem[®] (diltiazem HOL) □ 60 mg □ 90 mg □ 120 mg Sig: tid

Angina, arrhythmia, AV block (first degree), AV block (second or third degree — see conduction warning), bradycar-Cardiovascular.

dia, congestive heart failure, flushing, hypotension, palpitations, syncope.

Amnesia, gait abnormality, hallucina-

Nervous System: tions, insomnia, nervousness, paresthe

sia, personality change, somnolence, tinnitus, tremor.

Gastrointestinal: Anorexia, constipation, diarrhea, dysgeusia, dyspepsia, mild elevations of alkaline phosphatase, SGOT, SGPT, and

LDH (see hepatic warnings), vomiting, weight increase.

Dermatologic: Petechiae, pruritus, photosensitivity,

urticaria.

Amblyopia, dyspnea, epistaxis, eye irritation, hyperglycemia, nasal conges-Other-

tion, nocturia, osteoarticular pain, polyuria, sexual difficulties.

The following postmarketing events have been reported infrequently in patients receiving CARDIZEM. alopscia, infrequently in patients the control of the patients and the control of the patients. gingival hyperplasia, erythema multiforme, and leukopenia However, a definitive cause and effect between these events and CARDIZEM therapy is yet to be established.

See complete Professional Use Information before prescribing.

References: 1. Schroeder JS: Mod Med 1982;50(Sept):94-116. 2. Cohn PF, Braunwald E: Chronic ischemic heart disease, in Brounwald E (ed). <u>Heart Disease</u>. A <u>Textbook of Cardiovascular Medicine</u>, ed 2. <u>Philadelphia</u>, WB Saunders Co, 1984, chap 39. **3.** O'Rourke RA: <u>Am J Cardiol</u> 1985,56:34H-40H. **4.** McCall D, Walsh RA, Frohlich ED, et al: <u>Curr Probl Cardiol</u> 1985; 10(8):6-80. **5.** Frishman WH, Charlop S, Goldberger J, et al: Am J Cardiol 1985;56:41H-46H. & Shopiro W: Consultant 1984;24(Dec): 150-159 7. O'Hara MJ, Khurmi NS, Bowles MJ, et al: Am J Cardiol 1984;54:477-481. & Strauss WE, McIntyre KM, Parisi ĀF, et al: <u>Am J Cardiol</u> 1982; 49:560-566. **9.** Feldman RL, Pepine CJ, Whittle J, et al: <u>Am J Cardiol</u> 1982;49:554-559.



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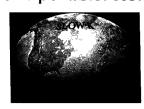
Capsule or tablet slow-release potassium chloride preparations should be reserved for patients who cannot tolerate, refuse to take, or have compliance problems with liquid or effervescent potassium preparations because of reports of intestinal and gastric ulceration and bleeding with slow-release KCI preparations.

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For patients who can't or won't tolerate liquid KCI.

*The most common adverse reactions to potassium salts are gastrointestinal side effects.

†Pooled mean serum potassium following oral administration of 30 mEq K-Tab compared to 24 mEq Slow-K in diuretic-treated hypertensives (n = 20) over 8 weeks.

CIBA

References: 1. Data on file. CIBA Pharmaceutical Company. 2. Skoutakis VA, Acchiardo SR, Wojciechowski NJ, et al: Liquid and solid potassium chloride: Bioavailability and safety. Pharmacotherapy 1980;4(6):392-397.

Skoutakis VA, Carfer CA, Acchiardo SR: Therapeutic assessment of Slow-K and K-Tab potassium chloride formulations in hypertensive patients treated with thiazide diuretics. Drug Intell Clin Pharm 1987;21:436-440.

Slow-K° potassium chloride USP Slow-Release Tablets 8 mEq (600 mg)

BRIEF SUMMARY (FOR FULL PRESCRIBING INFORMATION SEE PACKAGE INSERT)

NDICATIONS AND USAGE
BECAUSE OF REPORTS OF INTESTINAL AND GASTRIC ULCERATION AND
BLEEDING WITH SLOW-RELEASE POTASSIUM CHLORIDE PREPARATIONS, THESE DRIGS SHOULD BE RESERVED FOR THOSE PATIENTS
WHO CANNOT TOLERATE OR REFUSE TO TAKE LIQUID OR EFFERVES.
CENT POTASSIUM PREPARATIONS OR FOR PATIENTS IN WHOM THERE
IS A PROBLEM OF COMPLIANCE WITH THESE PREPARATIONS.

1. For therapeutic use in patients with hypokalemia with or without metabolic alkalosis; in digitalis intoxication and in patients with hypokalemic
samilial periodic paralysis.

2. For prevention of potassium depletion when the dietary intake of potassium is inadequate in the following conditions: patients receiving digitalis
and diuretics for congestive heart failure; hepatic cirrhosis with ascites;
states of aldosterone excess with normal renal function; potassium-losing
nephropathy; and certain diarrheal states.

3. The use of potassium stalts in patients receiving diuretics for uncomplicated essential hypertension is often unnecessary when such patients have
a normal dietary pattern. Serum potassium should be checked periodically,
however, and if hypokalemia occurs, dietary supplementation with potassium-containing foods may be adequate to control milder cases. In more
severe cases supplementation with potassium salts may be indicated.
CONTRANDICATIONS.
Potassium supplements are contraindicated in patients with hyperkalemia,
since a further increase in serum potassium salts may be indicated.
CONTRANDICATIONS.

All solid dosage forms of potassium supplements are sont produce cardiac arrest. Hyperkalemia may complicate any of the following conditions: chronic renal failure, systemic acidosis, acute dehydration, extensive tissue breakdown as in severe burns,
adrenal insufficiency, or the administration of a potassium-sparing diuretic
(e.g., spironolactone, triamterene) (see OVERIOSAGE).

All solid dosage forms of potassium supplements are contraindicated in
any patient in whom there is cause for arrest or delay in tablet passage
through the gast

WARNINGS
Hyperkalemia (See OVERDOSAGE).
Hyperkalemia (See OVERDOSAGE).
In patients with impaired mechanisms for excreting potassium, the administration of potassium salts can produce hyperkalemia and cardiac arrest. This occurs most commonly in patients given potassium by the intravous route but may also occur in patients given potassium orally. Potentially tatal hyperkalemia can develop rapidly and be asymptomatic.
The use of potassium salts in patients with chronic renal disease, or any other condition which impairs potassium excretion, requires particularly careful monitoring of the serum potassium concentration and appropriate dosage adjustment.

Interaction With Potassium-Sparing Diuretics
Hypokalemia should not be treated by the concomitant administration of
potassium sats and a potassium-sparing diuretic (e.g., spironolactone or
triamterene), since the simultaneous administration of these agents can
produce severe hopershapping.

poussion saits and a poussion-spanng outere (e.g., spironolactore or irramterene), since the simultaneous administration of these agents can produce severe hyperkalemia.

Gastrointestinal Lesions

Potassium chloride tablets have produced stenotic and/or ulcerative lesions

of the small bowel and deaths. These lesions are caused by a high localized concentration of potassium ion in the region of a rapidly dissolving tablet, which injures the bowel wall and thereby produces obstruction, hemorinage, or perforation. Slow-K is a wax-matrix tablet formulated to provide a controlled rate of release of potassium chloride and thus to minimize the possibility of a high local concentration of potassium on near the bowel wall. While the reported frequency of small-bowel lesions is much less with wax-matrix tablets (eas-tablets) and the size of the provide and the size of the size of the provide and the size of the si

ration considered it severe vomiting, abdominal pain, distention, or gastro-intestinal bleeding occurs.

Metabolic Acidosis
Hypokalemia in patients with metabolic acidosis should be treated with an alkalinizing potassium salt such as potassium bicarbonate, potassium ci-trate, or potassium acetate.

PRECAUTIONS

General:

The diagnosis of potassium depletion is ordinarily made by demonstrating hypokalemia in a patient with a clinical history suggesting some cause for potassium depletion. In interpreting the serum potassium incl., the physician should bear in mind that acute alkalosis per se can produce hypokalemain in the absence of a deficit in total body potassium, while acute arinsiss per se can increase the serum potassium concentration into the normal range even in the presence of a reduced total body potassium. Information for Patients

Indigres even in the presence of a decudent dual outp ploadsanin.

Indigration for Patients

Physicians should consider reminding the patient of the following:

To take each dose without crushing, chewing, or sucking the tablets.

To take this medicine only as directed. This is especially important if the patient is also taking both diruretics and digitalis preparations.

To check with the physician if there is trouble swallowing tablets or if the tablets seem to stick in the throat.

To check with the doctor at once if tarry stools or other evidence of gastrointestinal bleeding is noticed.

Laboratory Tests

Regular serum potassium determinations are recommended. In addition, during the treatment of potassium depletion, careful attention should be paid to acid-base balance, other serum electrolyte levels, the electrocardiogram, and the clinical status of the patient, particularly in the presence of cardiac disease, renal disease, or acidosis.

Drug Interactions

Cardiac disease, renal disease, or acidosis.

Drug Interactions
Possium-sparing diuretics: see WARNINGS.

Carcinogenesis, Mutagenesis, Impairment of Fertility
Long-term carcinogenicity studies in animals have not been performed.

Pregnancy Category C
Animal reproduction studies have not been conducted with Slow-K. It is also not known whether Slow-K can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. Slow-K should be given to a pregnant woman only if clearly needed.

Nursing Mothers
The normal potassium ion content of human milk is about 13 mEq/L. It is not known if Slow-K has an effect on this content. Caution should be exercised when Slow-K is administered to a nursing woman.

Safety and effectiveness in children have not been established.

ADVERSE REACTIONS
One of the management of the control of the

Pediatric Use
Safety and effectiveness in children have not been established.

ADVERSE REACTIONS
One of the most severe adverse effects is hyperkalemia (see CONTRAINDI-CATIONS, WARNINGS, and OVERDOSAGE). There also have been reports of upper and lower gastrointestinal conditions including obstruction, bleeding, ulceration, and perforation (see CONTRAINDICATIONS and WARNINGS); other factors known to be associated with such conditions were present in many of these patients.

The most common adverse reactions to oral potassium salts are nausea, vomiting, abdominal discomfort, and diarrhea. These symptoms are due to irritation of the gastrointestinal tract and are best managed by taking the dose with meals or reducing the dose.

Skin rash has been reported rarely.

OVERDOSAGE

The administration of oral potassium salts to persons with normal excretory mechanisms for potassium rarely causes serious hyperkalemia. However, if excretory mechanisms are impaired or if potassium is administered too rapidly intravenously, potentially statel hyperkalemia can result (see CON-TRAINDICATIONS and WARNINGS).) It is important to recognize that hyperalemia is usually asymptomatic and may be manifested only by an increased serum potassium concentration (6.5-8.0 mEq/L) and characteristic electrocardiographic changes (peaking of T waves, loss of P wave, depression of S-T segment, and prolongation of the Q-T interval). Later manifestations include muscle paralysis and cardiovascular collapse from cardiac arrest (9-12 mEq/L).

Treatment measures for hyperkalemia include the following: (1) elimination of foods and medications containing potassium and of potassium-paging diureties; (2) intravenous administration of 300-500 ml/hr of 10% dextrose solution containing notates and cardiovascular collapse from cardiac arrests. In treating the pyperkalemia in patients who have been stabilized on digitalis toxicity.

DOSAGE AND ADMINISTRATION

digitalis toxicity.

DOSAGE AND ADMINISTRATION

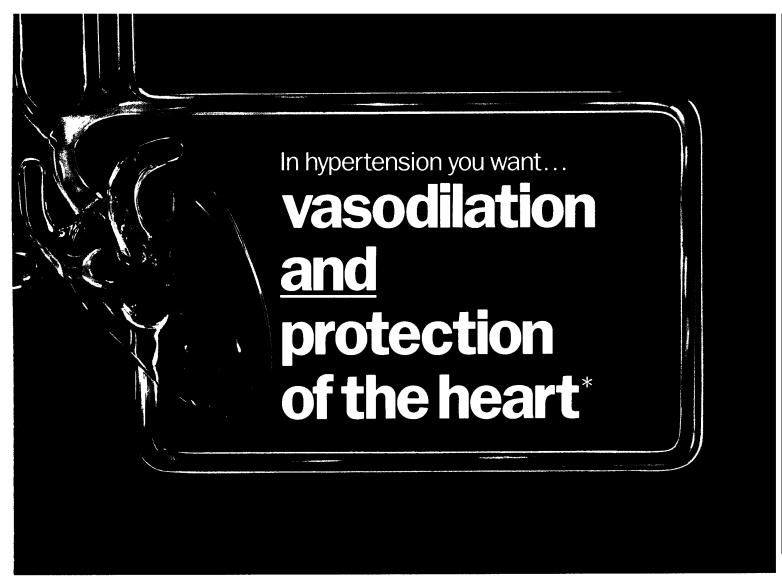
Dispense in tight, light-resistant container (USP).

Dist. by: CIBA Pharmaceutical Company Division of CIBA-GEIGY Corporation Summit, New Jersey 07901

C87-31 (Rev. 8/87)

CIBA

128-3568-A



NORMODYNE® gives you both (labetalol HCI) Tablets

	vasodilation	beta blockade
NORMODYNE (labetalol HCI) Tablets	V	V
Beta Blockers		
ACE Inhibitors		
Calcium Channel Blockers		

- ☐ Low incidence of impotence, fatigue, or cold extremities[†]
- ☐ Lipid and potassium levels are not adversely affected
- ☐ Minimizes risk of reflex tachycardia
- ☐ Maintains cardiac output
- ☐ Maintains exercise capacity
- $\hfill\square$ Does not adversely affect heart rate
- ☐ Maintains blood flow to vital organs
- ☐ Renal function is unimpaired

the most common side effects noted in treating mild to moderate hypertension with NORMODYNE (labetalol HCl) Tablets include dizziness (11%), nausea (6%), and fatigue (5%). Dyspepsia (3%), nasal stuffiness (3%), impotence (1%), and drows-

iness (<1%) occurred to a lesser degree. Overall, reports of symptomatic postural hypotension have been uncommon and have included rare instances of syncope. For complete side effects profile, see Prescribing Information.

For Brief Summary, please see reverse side of page.

Key Pharmaceuticals, Inc. Kenilworth, NJ 07033
World leader in drug delivery systems.

^{*}reduces double product (HR \times SBP) and minimizes the risk of reflex tachycardia

[†]Most adverse effects are mild, transient, and occur early in the course of treatment. In controlled clinical trials of three to four months' duration,

13071942-JBS PRODUCT INFORMATION

NORMODYNE® ed of labetaloi hydrochloride

Tablets BRIEF SUMMARY

INDICATIONS AND USAGE
NORMODYNE (labetalol HCI) Tablets are indicated in the management of hypertension. NORMODYNE Tablets may be used alone or in combination with other antihypertensive agents, especially thiazide

CONTRAINDICATIONS
NORMODYNE (labetalol HCl) Tablets are contraindicated in bronchial asthma, overt cardiac failure, greater than first degree heart block, cardiogenic shock, and severe bradycardia (see WARNINGS).

WARNINGS

Cardiac Failure Sympathetic stimulation is a vital component supporting circulatory function in congestive heart failure. Beta blockade carries a potential hazard of further depressing myocardial contractility and precipitating more severe failure. Although beta-blockers should be avoided in overt congestive heart failure, if necessary, labetalol HCl can be used with caution in patients with a history of heart failure who are well-compensated. Congestive heart failure, if necessary, labetalol HCl can be used with caution in patients with a history of heart failure who are well-compensated. Congestive heart failure has been observed in patients receiving labetalol HCl. Labetalol HCl does not abolish the inotropic action of digitalis on heart muscle.

In Pasients Without a History of Cardiac Failure in patients with latent cardiac insufficiency, continued depression of the myocardium with beta-blocking agents over a period of time can, in some cases, lead to cardiac failure. At the first sign or symptom of impending cardiac failure, patients should be fully digitalized and/or be given a diuretic, and the response observed closely. If cardiac failure continues, despite adequate digitalization and diuretic, NORMODYNE (labetalol HCl) therapy should be withdrawn (gradually if possible).

Exacerbation of Ischemic Heart Disease Following Abrupt Withdrawal Angina pectors is has not been reported upon labetalol HCl discontinuation. However, hypersensitivity to catecholamines has been observed in patients withdrawn from beta-blocker therapy; exacerbation of angina and, in some cases, myocardial infarction have occurred after abrupt discontinuation of such therapy. When discontinuing chronically administered NORMODYNE (labetalol HCl) particularly in patients with sichemic heart disease, the dosage should be gradually reduced over a period of one to two weeks and the patient should be carefully monitored. If angina markedly worsens or acute coronary insufficiency develops, NORMODYNE (labetalol HCl) administration' ops, NORMODYNE (labetalol HCJ) administration should be reinstituted promptly, at least temporarily, and other measures appropriate for the management of unstable angina should be taken. Patients should be warned against interruption or discontinuation of therapy without the physician's advice. Because coronary arrery disease is common and may be unrecognized, it may be prudent not to discontinue NORMODYNE (labetalol HCJ) therapy abruptly even in patients

mon and may be unrecognized, it may be prudent not to discontinue NORMODYNE (labetalol HCl) therapy abruptly even in patients treated only for hypertension.

Nonallergic Bronchospasm (e.g., chronic bronchitis and emphysema) Patients with bronchospastic disease should, in general, not receive beta-blockers. NORMODYNE may be used with caution, however, in patients who do not respond to, or cannot tolerate, other antihypertensive agents. It is prudent, if NORMODYNE is used, to use the smallest effective dose, so that inhibition of endogenous or exogenous beta-agonists is minimized.

Pheochromocytoma Labetalol HCl has been shown to be effective in lowering the blood pressure and relieving symptoms in patients with pheochromocytoma. However, paradoxical hypertensive responses have been reported in a few patients with this tumor; therefore, use caution when administering labetalol HCl to patients with pheochromocytoma.

Diabetes Mellius and Hypoglycemia Beta-adrenergic blockade may prevent the appearance of premonitory signs and symptoms (e.g., tachycardia) of acute hypoglycemia. This is especially important with labile diabetics. Beta-blockade also reduces the release of insulin in response to hyperglycemia; it may therefore be necessary to adjust the dose of antidiabetic drugs.

Major Surgery The necessity or desirability of withdrawing beta-blocking therapy prior to major surgery is controversial-Portracted severe hypotension and difficulty in restarting or maintaining a heart beat have been reported with beta-blockers. The effect of labetalol HCl's alpha-adnenergic activity has not been evaluated in this setting.

A synergism between labetalol HCl and halothane anesthesia has been shown (see Drug Interactions).

PRECAUTIONS

PRECAUTIONS
General Impaired Hepatic Function NORMODYNE (labetalol HCI)
Tablets should be used with caution in patients with impaired hepatic function since metabolism of the drug may be diminished.
Jaundice or Hepatic Dysfunction On rare occasions, labetalol HCl has been associated with jaundice (both hepatic and cholestatic). It is therefore recommended that treatment with labetalol HCl be stopped immediately should a patient develop jaundice or laboratory evidence of livet injury. Both have been shown to be reversible on stopping therapy.
Information for Patients.

liver injury. Both have been shown to be reversible on stopping therapy. Information for Patients
As with all drugs with beta-blocking activity, certain advice to patients being treated with labetalol HCl is warranted. This information is intended to aid in the safe and effective use of this medication. It is not a disclosure of all possible adverse or intended effects. While no incident of the abrupt withdrawal phenomenon (exacerbation of anging pectoris) has been reported with labetalol HCl, dosing with NORMODYNE Tablets should not be interrupted or discontinued without a physician's advice. Patients being treated with NORMODYNE Tablets should consult a physician at any sign of impending cardiac failure. Also, transient scalp tingling may occur, usually when treatment with NORMODYNE Tablets is initiated (see ADVERSE REACTIONS).

Laboratory Tests

As with any new drug given over prolonged periods, laboratory parameters should be observed over regular intervals. In patients with concomitant illnesses, such as impaired renal function, appropriate tests should be done to monitor these conditions.

concomitant illnesses, such as impaired renal function, appropriate tests should be done to monitor these conditions.

Drug Interactions

In one survey, 2.3% of patients taking labetalol HCl in combination with tricyclic antidepressants experienced tremor as compared to 0.7% reported to occur with labetalol HCl alone. The contribution of each of the treatments to this adverse reaction is unknown but the possibility of a drug interaction cannot be excluded.

Drugs possessing beta-blocking properties can blunt the bronchodilator effect of beta-receptor agonist drugs in patients with bronchospasm, therefore, doses greater than the normal anti-asthmatic dose of beta-agonist bronchodilator drugs may be required.

Cimetidine has been shown to increase the bioavailability of labetalol HCl. Since this could be explained either by enhanced absorption or by an alteration of hepatic metabolism of labetalol HCl, special care should be used in establishing the dose required for blood pressure control in such patients.

Synergism has been shown between halothane anesthesia and intravenously administered labetalol HCl. During controlled hypotensive anesthesia using labetalol HCl in association with halothane, high concentrations (3% or above) of halothane should not be used because the degree of hypotension will be increased and because of the possibility of a large reduction in cardiac output and an increase in central venous pressure. The anesthesiologist should be informed when a patient is receiving labetalol HCl.

Labetalol HCl blunts the reflex tachycardia produced by nitroglycerin without preventing its hypotensive effect. If labetalol HCl is used with nitroglycerin in patients with angina pectoris, additional antihypertensive effects may occur.

Drug/Laboratory Test Interactions
The presence of a metabolite of labetalol in the urine may result in falsely increased levels of urinary catecholamines when measured by a nonspecific trihydroxyindole (THI) reaction. In screening patients suspected of having a pheochromocytoma and being treated with labetalol HCl, specific radioenzymatic or high performance liquid chromatography assay techniques should be used to determine levels of catecholamines or their metabolites.

Carcinogenesis, Mutagenesis, Impairment of Fertility
Long-term oral dosing studies with labetalol HCl for 18 months in mice and for 2 years in rats showed no evidence of carcinogenesis.

Studies with labetalol HCl, using dominant lethal assays in rats and mice, and exposing microorganisms according to modified Ames tests, showed no evidence of mutagenesis.

potential risk to the ferus.

Nonteratogenic Effects
Infants of mothers who were treated with labetalol HCl for hypertension during pregnancy did not appear to be adversely affected by the drug. Oral administration of labetalol to rats during late gestation through weaning at doses of 2 to 4 times the MRHD caused a decrease in neonatal survival.

Labor and Delivery
Labor and Delivery
Labetalol HCl given to pregnant women with hypertension did not appear to affect the usual course of labor and delivery.

Nursing Mothers
Small amounts of labetalol (approximately 0.004% of the maternal does) are excreted in human milk. Caution should be exercised when NORMODYNE Tablets are administered to a nursing woman.

Pediatric Use Safety and effectiveness in children have not been established.

ADVERSE REACTIONS

Most adverse effects are mild, transient and occur early in the course of treatment. In controlled clinical trials of 3 to 4 months duration, discontinuation of NORMODYNE (labetalol HCl) Tablets due to one or more adverse effects was required in 7% of all patients. In these same trials, beta-blocker control agents led to discontinuation in 8 to 10% of patients, and a centrally acting alpha-agonist in 30% of patients.

The incidence rates of adverse reactions listed in the following table were derived from multicenter controlled clinical trials, comparing labetalol HCl, placebo, metopolo land proparalol, over treatment periods of 3 and 4 months. Where the frequency of adverse effects for labetalol HCl and placebo is similar, causal relationship is uncertain. The rates are based on adverse reactions considered probably drug-related by the investigator. If all reports are considered, the rates are somewhat higher (e.g., dizziness 20%, nausea 14%, fatigue 11%), but the overall conclusions are unchanged.

	Labetalol HCI (N = 227) %	Placebo (N = 98) %	Propranolol (N = 84) %	Metoprolo (N = 49) %
Body as a whole				
fatigue	5	0	12	12
asthenia	1	1	1	0
headache	2	1	1	2
Gastrointestinal				
nausea	_6	1	1	2
vomiting	< 1	Ó	o	ō
dyspepsia	3	1	1	0
abdominal pain	_0	0	1 2 0	0 0 2 0 0
diarrhea	<1	0	2	Ŏ
taste distortion	1	•	U	U
Central and Peripheral	Nervous Syst	ems		
dizziness)i	3	4	4
paresthesias	<u>></u> !	0	0	ŭ
drowsiness	< 1	2	2	2
Autonomic Nervous S	ystem		•	
nasal stuffiness	3	Ŏ	Õ	0
ejaculation failure	2	0	0	0 3 0
impotence	_¦	Ň	0	,
increased sweating	< 1	U	U	U
Cardiovascular		•	^	^
edema	. !	0	0	0
postural hypotension	1 1	0	0	12
bradycardia	U	U	,	12
Respiratory	•	0		2
dyspnea	2	U	1	2
Skin			•	
rash	1	0	0	0
Special Senses		_	_	_
vision abnormality	1	0	0	Ŏ
vertigo	2	1	0	0

The adverse effects were reported spontaneously and are representative of the incidence of adverse effects that may be observed in a properly selected hypertensive patient population, i.e., a group excluding patients with bronchospastic disease, over congestive heart failure, or other contrandications to beta-blocker therapy.

Clinical trials also included studies utilizing daily doses up to 2400 mg in more severely hypertensive patients. Certain of the side effects increased with increasing dose as shown in the table below which depicts the entire U.S. therapeutic trials data base for adverse reactions that are clearly or possibly dose related.

Labetalol HCl					
Daily Dose (mg)		200	300	400	600
Number of					
Patients		522	181	606	608
Dizziness (%)		2	3	3	3
Fatigue		2	1	4	- 4
Nausea		<1	0	1	. 2
Vomiting		0	0	<1	<1
Dyspepsia		1	0	2	1
Paresthesias		2	0	2	2
Nasal Stuffiness		1	1	2	2
Ejaculation Failure		0	2	1	2
Impotence		1	1	1	1
Edema		1	0	1	1
Labetalol HCl					
Daily Dose (mg)	800	900	1200	1600	2400
Number of					
Patients	503	117	411	242	175
Dizziness (%)	5	1	9	13	16
Fatigue	5	3	7	6	10
Nausea	4	0	7	11	19
Vomiting	< i	0	1	2	3
Dyspensia	1	0	2	2	4

Labetalol HCl Daily Dose (mg) (cont.)	800	900	1200	1600	2400
Paresthesias	1	1	2	5	5
Nasal Stuffiness	ż	į	4	Ś	6
Eiaculation Failure	3	õ	4	3	Š
Impotence	2	4	j	4	3
Edema	ī	Ó	ì	ż	2

Edema 1 0 1 2 2

In addition, a number of other less common adverse events have been reported in clinical trials or the literature:
Cardiovascular Postural hypotension, including rarely, syncope.
Central and Peripheral Nervous Systems Paresthesias, most frequently described as scalp tingling. In most cases, it was mild, transient and usually occurred at the beginning of treatment.
Collagen Disorders Systemic lupus erythematosus; positive antinuclear factor (ANF).
Eyes Dry eyes.
Immunological System Antimitochondrial antibodies.
Liver and Bildary System Cholestasis with or without jaundice.
Musculo-Sheletal System Muscle cramps; toxic myopathy.
Respiratory System Bonchospasm.
System Difficulty uricarial; bullous lichen planus; psoriaform; facial erythema; Peyronie's disease; reversible alopecia.
Urimary System Difficulty in micturition, including acute urinary bladder retention.
Following approval for marketing in the United Kingdom, a

Urinary System Difficulty in micrurition, including acute urinary bladder retention.

Following approval for marketing in the United Kingdom, a monitored release survey involving approximately 6,800 patients was conducted for further safety and efficacy evaluation of this product. Results of this survey indicate that the type, severity, and incidence of adverse effects were comparable to those cited above.

Potential Adverse Effects
In addition, other adverse effects not listed above have been reported with other beta-adrenergic blocking agents.

Central Nervous System Reversible mental depression progressing to catatonia; an acute reversible syndrome characterized by disorientation of time and place, short-term memory loss, emotional lability, slightly clouded sensorium, and decreased performance on neuropsychometrics.

Cardiouscular Intensification of AV block. See

CONTRAINDICATIONS.

Allergic Fever combined with aching and sore throat; laryngospasm; respiratory distress.

Hematologic Agranulocytosis; thrombocytopenic or nonthrom-bocytopenic purpura.

Gastroinustinal Mesenteric artery thrombosis; ischemic colitis. The oculomucocutaneous syndrome associated with the beta-blocker practool has not been reported with labetalol HCl.

Clinical laboratory tests: There have been reversible increases of serum transaminases in 4% of patients treated with labetalol HCl and tested, and more rarely, reversible increases in blood urea.

OVERDOSAGE

OVERDOSAGE

Overdosage with NORMODYNE (labetalol HCl) Tablets causes excessive brodycardia. Patients should be laid supine and their legs raised if necessary to improve the blood supply to the brain. The following additional measures should be employed if necessary: Excessive brodycardia—administer atropine (3.0 mg). If there is no response to vagal blockade, administer atropine (3.0 mg). If there is no response to vagal blockade, administer atoproterenol cautiously. Cardiac fallare — administer adigitalis glycoside and a diuretic. Hypotension — administer vasopressors, e.g., norepinephrine. There is pharmacological evidence that norepinephrime may be the drug of choice. Bronchospasm — administer a betag-stimulating agent and/or a theophylline preparation.

Gastric lavage or pharmacologically induced emesis (using syrup of ipseca() is useful for removal of the drug shortly after ingestion. Labetalol HCl can be removed from the general circulation by hemodialysis.

The oral LDsy value of labetalol HCl in the mouse is approximately 600 mg/kg and in the rat is greater than 2 gm/kg. The intravenous LDso in these species is 50 to 00 mg/kg.

in these species is 50 to 60 mg/kg.

DOSAGE AND ADMINISTRATION
DOSAGE MUST BE INDIVIDUALIZED. The recommended initial dose is 100 mg rwice daily whether used alone or added to a diuretic regimen. After 2 or 3 days, using standing blood pressure as an indicator, dosage may be titrated in increments of 100 mg bid every 2 or 3 days. The usual maintenance dosage of laberalol HCl is between 200 and 400 mg rwige daily.

Since the full antihypertensive effect of labetalol HCl is usually seen within the first one to three hours of the initial dose or dose increment, the assurance of a lack of an exaggerated hypotensive response can be clinically established in the office setting. The antihypertensive effects of continued dosing can be measured at subsequent visits, approximately 12 hours after a dose, to determine whether further titration is necessary. Patients with severe hypertension may require from 1200 mg to 2400 mg per day, with or without thiazide diuretics. Should side effects (principally nausea or dizziness) occur with these doses administered bid, the same total daily dose administered there times daily may improve tolerability and facilitate further titration. Titration increments should not exceed 200 mg twice daily.

When a diuretic is added, an additive antihypertensive effect can be expected. In some cases this may necessitate a labetalol HCl dosage adjustment. As with most antihypertensive drugs, optimal dosages of NORMODYNE Tablets are usually lower in patients also receiving a diuretic.

diuretic.

When transferring patients from other antihypertensive drugs,
NORMODYNE Tablets should be introduced as recommended and the dosage of the existing therapy progressively decreased.

HOWNOTINE is alabets should be introduced as recommended and the dosage of the existing therapy progressively decreased.

HOW SUPPLIED

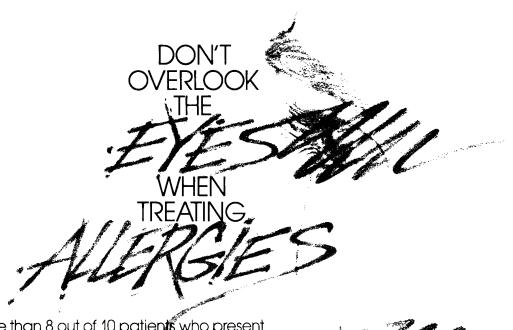
NORMODYNE (labetalol HCl) Tablets, 100 mg, light-brown, round, scored, film-coated tablets engraved on one side with Schering and product identification numbers 244, and on the other side the number 100 for the strength and "NORMODYNE"; bottles of 100 (NDC-0085-0214-04), 500 (NDC-0085-0214-05), and box of 100 for unit-dose dispensing (NDC-0085-0244-08).

NORMODYNE (labetalol HCl) Tablets, 200 mg, white, round, scored, film-coated tablets engraved on one side with Schering and product identification numbers 752, and on the other side the number 200 for the strength and "NORMODYNE"; bottles of 100 (NDC-0085-0752-04), 500 (NDC-0085-0752-05), box of 100 for unit-dose dispensing (NDC-0085-0752-08), and Parient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0752-03).

NORMODYNE (labetalol HCl) Tablets, 300 mg, blue, round, film-coated tablets engraved on one side with Schering and product identification numbers 438, and on the other side the number 300 for the strength and "NORMODYNE"; bottles of 100 (NDC-0085-0438-03), 500 (NDC-0085-0438-04), and Parient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-04), and Parient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-04), and Parient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-05), and Parient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56 (4 bottles of 14 tablets) (NDC-0085-0438-06), and Patient Calendar Package of 56

Key Pharmaceuticals, Inc. Kenilworth, NJ 07033 USA

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More than 8 out of 10 patients who present with allergic rhinitis may suffer from concurrent allergic ocular signs and symptoms —itchy, scratchy eyes, erythema and edema, tearing, irritation—according to two recent studies.¹²

What's worse is that many patients don't mention their ocular symptoms when reporting allergic rhinitis unless specifically asked.

So look for the overlapping clinical symptoms and then confidently treat alleraic ocular disorders* with OPTICROM. It has a proven clinical record of efficacy with freedom from serious side effects or ocular toxicity.



THE SOLUTION FOR ALLERGIC OCULAR DISORDERS

Reference: 1,2. Data on file, Fisons Corporation. Independent Studies by DTW Market Research Group. July 1985

*See below for listing of certain allergic ocular disorders

INDICATIONS AND USAGE: OPTICROM is indicated in the treatment of certain allergic ocular disorders referred to by the terms vernal keratoconjunctivitis, vernal conjunctivitis, glant papillary conjunctivitis, vernal keratitis, and allergic keratoconjunctivitis. The etiologic factors are unknown, but common airborne allergens and contact lenses have been implicated.

have been implicated."

Symptomatic response to therapy (decreased itching, tearing, redness, and discharge) is usually evident within a few days, but longer treatment for up to six weeks is sometimes required. Once symptomatic improvement has been established, therapy should be continued for as long as needed to sustain improvement.

If required, corticosteroids may be used concomitantly with OPTICROM.

Users of soft (hydrophilic) contact lenses should refrain from wearing lenses while under treatment with OPTICROM (see Contraindications). Wear can be resumed within a few hours after discontinuation of the drug.

CONTRAINDICATIONS: OPTICROM is contraindicated in those patients who have shown hypersensitivity to cromolyn sodium or to any of the other ingredients. As with all ophthalmic preparations containing benzalkonium chloride, patients are advised not to wear soft contact lenses during treatment with OPTICROM.

are advised not to wear soit contact lenses during treatment with OPTICROM.

PRECAUTIONS: General: Patients may experience a transient stinging or burning sensation following application of OPTICROM.

The recommended frequency of administration should not be exceeded. The dose for adults and children is 1-2 drops in each eye 4-6 times a day at regular intervals.

Carcinogenesis, Mutagenesis, and Impairment of Fertility: Long term studies in mice (12 months intraperitoneal treatment followed by six months observation), hamsters (12 months intraperitoneal treatment followed by 12 months observation), and rats (18 months subcutaneous treatment) showed no neoplastic effect of cromolyn sodium. No evidence of chromosomal damage or cytotoxicity was obtained in various mutagenesis studies.

mutagenesis studies

No evidence of impaired fertility was shown in laboratory animal reproduction studies.

Pregnancy: Pregnancy Category B. Reproduction studies with cromolyn sodium administered parenterally to pregnant mice. rats and rabbits in doses up to 338 times the human clinical doses produced no evidence of fetal malformations. Adverse fetal effects (increased resorption and decreased fetal weight) were noted only at the very high parenteral doses that produced maternal toxicity. There are, however, no adequate and well controlled studies in pregnant women. Because animal reproduction studies are not always predictive of human response, this drug should be used during pregnancy only if clearly needed.

Nursing Mothers: It is not known whether this drug is excreted in human milk, caution should be exercised when OPTICROM is administered to a nursing woman.

Pediatric Use: Safety and effectiveness in children below the age of 4 years have not been

established.

ADVERSE REACTIONS: The most frequently reported adverse reaction attributed to the use of OPTICROM, on the basis of reoccurrence following readministration, is transient ocular stinging or burning upon instillation.

The following adverse reactions have been reported as infrequent events. It is unclear whether they are attributable to the drug; conjunctival injection, watery eyes, itchy eyes, dryness around the eye, puffy eyes, eye irritation, styes.

CAUTION: Federal law prohibits dispensing without prescription.

REFERENCE: 1. Allansmith MR, Abelson MB. Ocular Allergies. In: *The Cornea*, ed. by G. Smolin, RA Thoft, Little, Brown and Co., Boston/Toronto, 1983: 231-43. See package insert for full prescribing information.

OPTICROM* is a registered trademark of Fisons plo Made in England FC7201 Revised 07/84 F1S-026 FISONS CORPORATION BEDFORD, MA 01730

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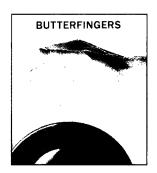
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without a history of allergy or bronchial asthma. Possible exacerbation or activation of systemic lupus erythematosus has been reported with thiazide diuretics.

Precautions: The bioavailability of the hydrochlorothiazide component of 'Dyazide' is about 50% of the bioavailability of the single entity. Theoretically, a patient transferred from the single entities of triamterene and hydrochlorothiazide may show an increase in blood pressure or fluid retention. Similarly, it is also possible that the lesser hydrochlorothiazide bioavailability could lead to increased serum potassium levels. However, extensive chinical experience with 'Dyazide' suggests that these conditions have not been commonly observed in clinical practice. Angiotensin-converting enzyme (ACE) inhibitors can elevate serum potassium; use with caution with 'Dyazide'. Do periodic serum electrolyte determinations (particularly important in patients vomiting excessively or receiving parenteral fluids, and during concurrent use with amphotericin B or corticoteroids or corticoteopin/ACTHI). Periodic BUN and serum creatinine determinations should be made, especially in the elderly, diabetics or those with suspected or confirmed renal insufficiency. Cumulative effects of the drug may develop in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with impaired hepatic function. They can precipitate coma in patients with severe liver disease. Observe regularly for possible blood dy failure have been reported in patients on 'Dyazide' when treated with indomethacin. Therefore, caution is advised in administering

nonsteroidal anti-inflammatory agents with 'Dyazide'.

* Not for initial therapy. See brief summary.

following may occur: transient elevated BUN or creatinine or both, hyperglycemia and glycosuria (diabetic insulin requirements may be attered), hyperuricemia and gout, digitalis intoxication (in hypokalemia), decreasing alkali reserve with possible metabolic acidosis. Dyazide interferes with fluorescent measurement of quinidine. Hypokalemia is uncommon with 'Dyazide', but should it develop, corrective measures should be taken such as potassium sinh syndromentation or increased dietary litaks of palassium sinh.

This drug is not indicated for initial therapy of edema or hypertension. Edema or hypertension. pertension. Edema or hypertension requires therapy ated to the individual. If this combination represents the dosage so determined, its use may be more convenient in patient management. Treatment of hypertension and edema is not static, but must be reevaluated as conditions in each

ontraindications: Concomitant use with other potassium-paring agents such as spironolactone or amiloride. Further use anuria, progressive renal or hepatic dysfunction, hyperkalemia, re-existing elevated serum potassium. Hypersensitivity to either omponent or other sulfonamide-derived drugs. Varnings: Do not use potassium supplements, dietary or varnings: Do not use potassium supplements, dietary or

reactions seen in adults. Thiazides appear and triamterene may appear in breast milk. If their use is essential, the patient should stop nursing. Adequate information on use in children is not available. Sensitivity reactions may occur in patients with or

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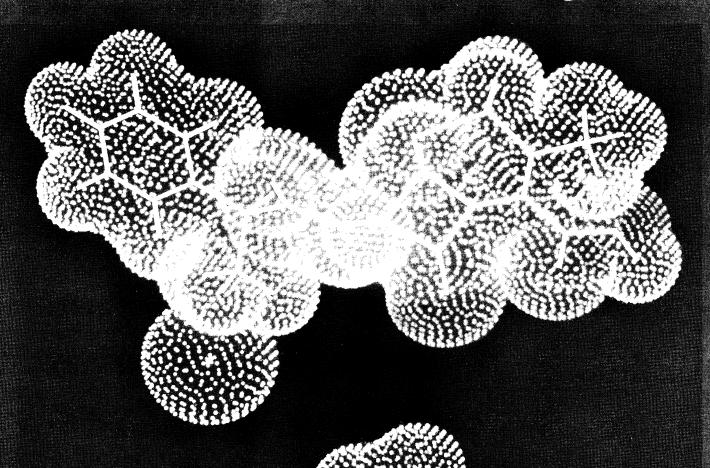


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Keftab is contraindicated in patients with known allergy to the cephalosporins and should be given cautiously to penicillinsensitive patients.

Penicillin is the drug of choice in the treatment and prevention of streptococcal infections, including the prophylaxis of rheumatic fever.

Due to susceptible strains of *Staphylococcus aureus* and or differency to straptococcu. Due to susceptible strains of *Eschorchia con Protectini trains* and Kingger and Due to susceptible strains of group Ald tiernoly to straptococci.

KEFTAB™

(cephalexin hydrochloride monohydrate)

Summary: Consult the package literature for prescribing information.

Indications and Usage:

Respiratory tract infections caused by susceptible strains of Streptococcus pneumoniae and group A β -hemolytic streptococci.

Skin and skin structure infections caused by susceptible strains of Staphylococcus aureus and/or β -hemolytic streptococci.

Bone infections caused by susceptible strains of S aureus and/or Proteus mirabilis.

Genitourinary tract infections, including acute prostatitis, caused by susceptible strains of Escherichia coli, P mirabilis, and Klebsiella sp.

Contraindication: Known allergy to cephalosporins.

Warnings: KEFTAB SHOULD BE ADMINISTERED CAUTIOUSLY TO PENICILLIN-SENSITIVE PATIENTS. PENICILLINS AND CEPHALOSPORINS SHOW PARTIAL CROSS-ALLERGENICITY. POSSIBLE REACTIONS INCLUDE ANAPHYLAXIS.

Administer cautiously to allergic patients.

Pseudomembranous colitis has been reported with virtually all broad-spectrum antibiotics. It must be considered in differential diagnosis of antibiotic-associated diarrhea. Colon flora is altered by broad-spectrum antibiotic treatment, possibly resulting in antibiotic-associated colitis.

Precautions:

- Discontinue Keftab in the event of allergic reactions to it.
- Prolonged use may result in overgrowth of nonsusceptible organisms.
- Positive direct Coombs' tests have been reported during treatment with cephalosporins.
- Keftab should be administered cautiously in the presence of markedly impaired renal function. Although dosage adjustments in moderate to severe renal impairment are usually not required, careful clinical observation and laboratory studies should be made.
- Broad-spectrum antibiotics should be prescribed with caution in individuals with a history of gastrointestinal disease, particularly colitis.
- Safety and effectiveness have not been determined in pregnancy and lactation. Cephalexin is excreted in mother's milk. Exercise caution in prescribing Keftab for these patients.
- Safety and effectiveness in children have not been established.

Adverse Reactions:

- Gastrointestinal, including diarrhea and, rarely, nausea and vomiting. Transient hepatitis and cholestatic jaundice have been reported rarely.
- Hypersensitivity in the form of rash, urticaria, angioedema, and, rarely, erythema multiforme, Stevens-Johnson syndrome, or toxic epidermal necrolysis.
- · Anaphylaxis has been reported.
- Other reactions have included genital/anal pruritus, genital moniliasis, vaginitis/vaginal discharge, dizziness, fatigue, headache, eosinophilia, neutropenia, and thrombocytopenia; reversible interstitial nephritis has been reported rarely.
- Cephalosporins have been implicated in triggering seizures, particularly in patients with renal impairment.
- Abnormalities in laboratory test results included slight elevations in aspartate aminotransferase (AST, SGOT) and alanine aminotransferase (ALT, SGPT). False-positive reactions for glucose in the urine may occur with Benedict's or Fehling's solution and Clinitest® tablets but not with Tes-Tape® (Glucose Enzymatic Test Strip, USP, Lilly).

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PHYSICIANS WANTED

PACIFIC NORTHWEST, NEAR EUGENE, OR-EGON. BE/BC Internist wanted to join Family Practice group. Share call with Internists. Minimum salary guarantee. Good schools/outdoor recreation; University of Oregon/cultural events within 30 minutes. Contact John Hoopes, Cottage Grove Hospital, 1340 Birch St, Cottage Grove, OR 97424; (503) 942-0511

NEAR STANFORD. Six Internists, all subspecialty trained and members of clinical faculty at Stanford, interested in an Associate with subspecialty interest and training. Should be well grounded in Internal Medicine. Send CV to Dr Bigler, El Camino Internal Medical Group, 125 South Dr, Mountain View, CA 94040.

IDAHO. Family Practitioner with interest in OB wanted to join three Family Practitioners serving scenic northern Idaho community. Hospital provides complete assistance-office, salary, full benefits. BE/BC and cesarean section experience required. Enjoy outdoor recreation, rural lifestyle. Contact Jean Erickson, PROSEARCH, 305 NE 102nd Ave, Portland, OR 97220-4199; (503) 256-2070.

SOUTH CENTRAL WASHINGTON COMMUNITY seeks BE/BC Internist for solo practice. Share office space with two other physicians. First year income guarantee and other assistance. Great income potential for right candidate! Progressive 38-bed hospital has CT services. Excellent schools and recreation. Contact PROSEARCH, 305 NE 102nd Ave, Portland, OR 97220; (503) 256-4488

CALIFORNIA CENTRAL VALLEY. Desire BC/BE Family Practitioner to join busy two man Family Practice. Easy drive to San Francisco or mountains. Growing area; city of 40,000; good schools. Guarantee plus benefits. Reply to Number 59, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602

NORTHERN CALIFORNIA OB/GYN. Beautiful Marin County, California OB/GYN practice looking for BC/BE female/male OB/GYN who is interested in joining a busy and successful practice of two MDs, one midwife, and four NPs. Progressive, busy OB practice offering family centered maternity care. Pro-choice philosophy. Opportunity for university affiliation. Lovely physical facilities with potential for investment. Must be well trained, enjoy working hard, committed to women's health care and fun to work with. Compensation package with partnership available in one year. Six weeks time off each year. Send résumé to WHMA/NEB, PO Box 1773, Ross, CA 94957.

PHYSICIANS WANTED

BC/BE INTERNIST to associate with General Surgeon, OB/GYN, Pediatrician, Internist, and three FPs in well-established rural practice. Send CV to R. F. LeBlond, MD, Park Clinic, Box 1139, Livingston, MT 59047

INTERNIST. Live in San Francisco and commute to nearby rural area for four two-night shifts per month in combined Internal Medicine/Emergency room practice. Four Internists currently working in stable group. Practice quality medicine in the country where you can make a greater impact and enjoy lots of free time wherever you like to live. 72k per year. Charles Rath, MD, 199 E. Webster St, Colusa, CA 95932; (916) 458-7739.

INTERNIST: SAN FRANCISCO. BC/BE Internist wanted to join primary care group in San Francisco. We are currently three doctors sharing call, wish a fourth to expand call schedule and purchase a well-established practice in our area from retiring Internist (former president of ASIM). Send résumé to John Pierce, MD, 3620 Army St, San Francisco, CA 94110, or call (415) 826-7577

GENERAL INTERNIST needed for large hospitalbased multispecialty clinic. University associated residency program. Attractive salary and complete benefit package. Pleasant setting. BC/BE. California license required. Contact Dennis L. Ostrem, MD, Chief Internal Medicine, The Permanente Medical Group, Inc, PO Box 254999, Sacramento. CA 95865-4999 or call (916) 973-5781. An Equal Opportunity Employer.

FAMILY PRACTICE PHYSICIANS. Immediate positions availabe for BE/BC Family Practice Physicians to join a growth-oriented southern California group with an expanding PCCM program. Daytime office practice only; no nights, weekends or Obstetrics. Full-time permanent positions with guaranteed incomes and paid malpractice. Send CV to Medical Director, PO Box 16027, Long Beach, CA 90806; or call (213) 590-9696.

NORTHERN SAN FRANCISCO BAY AREA: Seeking Physician BC/BE in Internal Medicine for Internist position in growing department. Kaiser Permanente Medical Center, 1550 Gateway Blvd, Fairfield, CA 94533; (707) 427-4200.

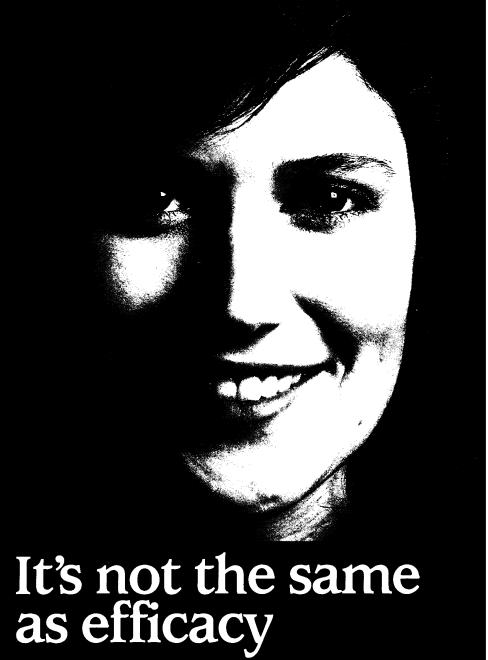
EMERGENCY GROUP seeking career oriented ACLS, ATLS physician for immediate opening. Moderate volume, income \$120,000. Great outdoor activities including fishing, boating, skiing and sailing in south central Washington. Send CV to KEP, PO Box 6192, Kennewick, WA 99336; or call (509) 627-1798.

(Continued on Page 252)

In the treatment of chronic anxiety...



Sedation...



BuSpar relieves anxiety and returns your patient to normal activity with no more sedation than induced by placebo¹...and without impairing psychomotor function in most patients*² or producing a benzodiazepine withdrawal syndrome upon discontinuation³

The first choice for chronic anxiety



^{*}Because the effects of BuSpar in any individual patient may not be predictable, patients should be cautioned about operating an automobile or using complex machinery until they are reasonably certain that BuSpar treatment does not affect them adversely. For Brief Summary, please see following page.

BuSpar (buspirone HCI)

References: 1. Newton RE, et al. A review of the side effect profile of buspirone. Am J Med 1996, 80(3B) 17-21. 2. Moskowitz H and Smiley A. Effects of chronically administered buspirone and diazepam on driving-related skills performance. J Clin Psychiatry 1982, 43(12, Sec 2), 45-55. 3. Lader M. Assessing the potential for buspirone dependence or abuse and effects of its withdrawal. Am J Med 1987:82(5A):20-2

Contraindications: Hypersensitivity to buspirone

Warnings: The administration of BuSpar to a patient taking a monoamine oxidase inhibitor (MAOI) may pose a hazard. Since blood pressure has become elevated when BuSpar was administered concomitantly with an MAOI, such concomitant use is not recommended. BuSpar should not be employed in ieu of appropriate antipsycholic treatment.

Precautions: General—Interference with cognitive and motor performance. Although buspirone is less sedating than other anxiolytics and does not produce significant functional impairment, its CNS effects in a given patient may not be predictable; therefore, patients should be cautioned about operating an automobile using complex machinery until they are reasonably certain that buspirone does not affect them adversely Although buspirone has not been shown to increase alcohol-induced impairment in motor and mental per formance, it is prudent to avoid concomitant use with alcohol.

Potential for withdrawal reactions in sedative/hypnotic/anxiolytic drug dependent patients: Because bus-

pirone will not block the withdrawal syndrome often seen with cessation of therapy with benzodiazepines and other common sedative/hypnotic drugs, before starting buspirone withdraw patients gradually from their prior treatment, especially those who used a CNS depressant chronically. Rebound or withdrawal symptoms may occur over varying time periods, depending in part on the type of drug and its elimination half-life. The withdrawal syndrome can appear as any combination of irritability, anxiety, agitation, insomnia, tremor, ab-dominal cramps, muscle cramps, vomiting, sweating, flu-like symptoms without fever, and occasionally,

Possible concerns related to buspirone's binding to dopamine receptors: Because buspirone can bind to central dopamine receptors, a question has been raised about its potential to cause acute and chronic changes in dopamine mediated neurological function (e.g., dystonia, pseudoparkinsonism, akathisia, and tardive dyskinesia). Clinical experience in controlled trials has failed to identify any significant neurolepticlike activity, however, a syndrome of restlessness, appearing shortly after initiation of treatment, has been reported; the syndrome may be due to increased central noradrenergic activity or may be attributable to

dopaminergic effects (ie. represent akathisia).

Information for Patients — Patients should be instructed to inform their physician about any medica tions, prescription or non-prescription, alcohol or drugs they are now taking or plan to take during treatment with buspirone; to inform their physician if they are pregnant, are planning to become pregnant, or become

pregnant while taking buspirone; to inform their physician if they are breast feeding; and not to drive a car or operate potentially dangerous machinery until they experience how this medication affects them. **Drug Interactions**—Concomitant use with other CNS active drugs should be approached with caution (see **Warnings**). Concomitant use with trazodone may have caused 3- to 6-fold elevations on SGPT (ALT) in

a few patients. Buspinne does not displace tightly bound drugs like phenytoin, propranolol, and warfarin from serum proteins, but may displace less firmly bound drugs like digoxin.

Carcinogenesis, Mutagenesis**, Impairment of Fertility**—No evidence of carcinogenic potential was observed in rats or mice: buspirone did not induce point mutations, nor was DNA damage observed, chromosomal aberrations or abnormalities did not occur.

Pregnancy: Teratogenic Effects—Pregnancy Category B: Should be used during pregnancy only if

Nursing Mathers—Administration to nursing women should be avoided if clinically possible.

Pediatric Use—The safety and effectiveness have not been determined in individuals below 18 years of age Use in the Elderly—No unusual, adverse, age-related phenomena have been identified in elderly patients receiving a total, modal daily dose of 15 mg.

Use in Patients with Impaired Hepatic or Renal Function—Since buspirone is metabolized by the

liver and excreted by the kidneys, it is not recommended in severe hepatic or renal impairment.

Adverse Reactions (See also Precautions): Commonly Observed—The more commonly observed unloward events include dizziness, nausea, headache, nervousness, lightheadedness, and excitement. Associated with Discontinuation of Treatment—The more common events causing discontinuation included: central nervous system disturbances (3.4%), primarily dizziness, insomnia, nervousness, drows-iness, lightheaded feeling, gastrointestinal disturbances (1.2%), primarily nausea, miscellaneous disturbances (1 1%), primarily headache and fatigue. In addition, 3 4% of patients had multiple complaints, none of which could be characterized as primary. *Incidence in Controlled Clinical Trials*—Adverse events reported by 1% or more of 477 patients who

Incidence in Committee Crimical Trials—Adverse Vents reported by 3 of more of 47 patients win received buspirone in four-week, controlled trials: Cardiovascular. Tachycardia/palpitations 1%. CNS: Diz-ziness 12%, drowsiness 10%, nervousness 5%, insomnia 3%, lightheadedness 3%, decreased concentra-tion 2%, excitement 2%, anger/hostility 2%, confusion 2%, depression 2%. EENT. Blurred vision 2%. Gastroinlestinal: Nausea 8%, dry mouth 3%, abdominal/gastric distress 2%, diarrhea 2%, constipation 1%, vomiting 1% Musculoskeletal: Musculoskeletal aches/pains 1% Neurological: Numbness 2%, paresthesia 1%, incoordination 1%, tremor 1%. Skin: Skin rash 1% Miscellaneous: Headache 6%, fatigue 4%, weak-

Other Events Observed During the Entire Pre-marketing Evaluation—The relative frequency of all other undestrable events reasonably associated with the use of buspirone in approximately 3000 subjects who took multiple doses of the drug under well-controlled, open, and uncontrolled conditions is defined as follows: Frequent are those occurring in at least 1700 patients: infrequent are those occurring in 1700 to 1700 patients. Cardiovascular requent, non-specific chest pain; infrequent: syncope, hypotension, hypertension, rare: cerebrovascular accident, congestive heart failure, myocardial infarction, cardiomyopathy, bradycardia. Central Nervous System—Irequent dema disturbances, infrequent depersonalization, dysphoria, noise infolerance, euphoria, akathisia, learfulness, loss of interest, disassociative reaction, hallucinations, suicidal ideation, seizures, rare: feelings of claustrophobia, cold intolerance, stupor, slurred speech, psychosis. EENT—Trequent: timitus, sore throat, nasal congestion, infrequent: redness and tiching of the eyes, altered taste, altered smell, conjunctivitis; rare: inner ear abnormatity, eye pain, photophobia, pressure on eyes. Endocrine—rare: galactorrhea, thyroida borormatity. Gastrointestinal—infrequent: flatulence, ancovai, increased appetite, salivation, irritable colon. Other Events Observed During the Entire Pre-marketing Evaluation—The relative frequency of normality. Gastrointestinal—infrequent: flatulence, anorexia, increased appetite, salivation, irritable colon, rectal bleeding, rare, burning of the tongue. Genitourinary—infrequent: urinary frequency, urinary hesitancy, menstrual irregularity and spotting, dysuria, rare, amenorthea, pelvic inflammatory disease, enuresis, nocturia. Musculoskeletal—infrequent: muscle cramps, muscle spasms, rigid/stiff muscles, arthratigias. Neurological—infrequent: involuntary movements, slowed reaction time, rare, muscle weakness. Respiratory infrequent: hyperventilation, shortness of breath, chest congestion, rare: epistaxis. Sexual Function—infrequent: decreased or increased libido, rare: delayed ejaculation, impotence. Skin—infrequent: deema pruntus. flushing, easy bruising, hair loss, dry skin, facial edema, blisters, rare, acne, thinning of nails. *Clinical Laboratory*—infrequent: increases in hepatic aminotransferases (SGOT, SGPT), rare: eosinophilia, leu-

ical Laboratory—infrequent: increases in hepatic aminotransterases (SGUI, SGP1), rare: eosinophilia, leukopenia, thrombocytopenia. Miscellaneous—infrequent: weight gain, fever, roaring sensation in the head, weight loss, malaise, rare: alcohol abuse, bleeding disturbance, loss of voice, hiccoughs.

Drug Abuse and Dependence: Controlled Substance Class—Not a controlled substance.

Physical and Psychological Dependence—Buspirone has shown no potential for abuse or diversion and there is no evidence that it causes tolerance, or either physical or psychological dependence. However, since it is difficult to predict from experiments the extent to which a CNS active drug will be misused, diverted, and/or abused once marketed, physicians should carefully evaluate patients for a history of drug abuse and follow such patients (lose), voice vigent them for sings of busingene misuse or abuse (n. devel-

verted, afford above once makeley, physicians should calerately evaluate patients for a mistary of unity abuse and follow such patients closely, observing them for signs of buspirone misuse or abuse (e.g., development of tolerance, incrementation of dose, drug-seeking behavior).

Overdosage: Signs and Symptoms—At doses approaching 375 mg/day the following symptoms were observed nauseax owniting, dizziness, drowsiness, miosis, and agastric distress. No deaths have been reported in humans either with deliberate or accidental overdosage.

Recommended Overdose Treatment—General symptomatic and supportive measures should be used along with immediate gastric lavage. No specific antidote is known and dialyzability of buspirone has not been determined

For complete details, see Prescribing Information or consult your Mead Johnson Pharmaceuticals Repre-

Mead THEM PHARMACEUTICALS

Bristol-Myers U.S. Pharmaceutical and Nutritional Group · Evansville, Indiana 47721 U.S.A. MJL 7-4223 ADS

RESULTS







CLASSIFIED INFO (415)882-5178

(Continued from Page 250)

PASUAS VANDO GASTROENTEROLOGIST-BC/BE to join two Gastroenterologists in busy private clinical practice located in highly desirable Los Angeles suburb. Strong clinical and endoscopic skills needed. We perform all diagnostic and therapeutic procedures including Laser and Sphincterotomy. Competitive salary and benefits with early partnership potential. Available July 1988 or sooner. Send CV to Number 80, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602

UNIVERSITY OF CALIFORNIA, Irvine, Department of Medicine is seeking a full-time faculty person as General Internist for expanding academic group practice. Combined fee-for-service/capitation. Duties include 80-90% clinical practice in multispecialty faculty clinic, 10-20% teaching residents and students ambulatory care and inpatient medicine. Division of General Internal Medicine with strong commitment to teaching, practice, and research. Competitive salary and benefits. Affirmative action/equal opportunity employer. Send CV to Jeremiah Tilles, MD, UCI, Department of Medicine, Route 81, 101 City Drive South, Orange, CA 92668

FAMILY PHYSICIANS, BC/BE, full- and part-time positions available with Obstetrics optional, to work with multispecialty group practice in the Seattle area. Attractive salary benefits. Contact Sharon Courlas, MD, (206) 326-4147. Send CV to Pacific Health Associates, 1200 12th Ave South, Seattle, WA 98144, Attn: Mary Anderson

BC/BE INTERNIST. In northern California wine country. Join two man group in private practice of Internal Medicine, Subspecialty interest OK. Reply to Number 81, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602

CRESCENT CITY, CALIFORNIA. Exciting position available at a growing 24,000-visit ER in a rural, coastal community. Fee-for-service with possibility of six figure income. Send CV to Art B. , Wong, MĎ, FACEP, 1 Maritime Plaza, Ste 710, San Francisco, CA 94111

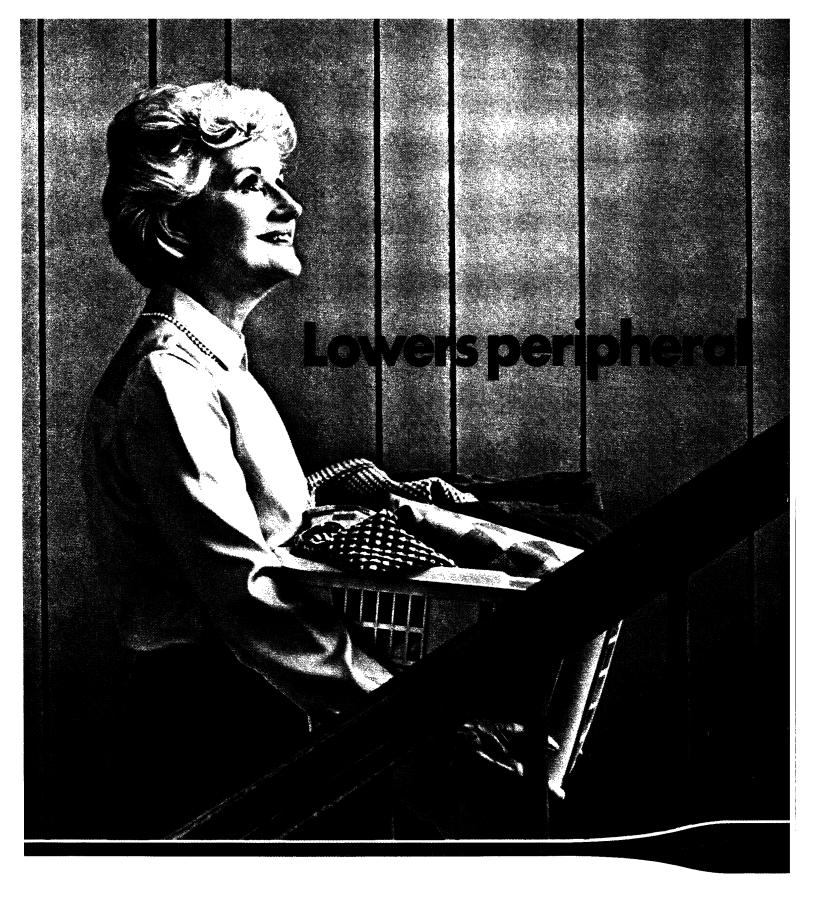
GENERAL INTERNIST, BC/BE to join group practice in smaller community near Modesto, California. Join four other Internists and two Physicians Assistants. Full range of hospital and office practice. Must be interested in Primary Care focus, but will consider sub-specialty in Cardiology or Infectious Diseases. May call (collect) or write to Suzanne Meyer, MD, 6449 3rd St, Riverbank, CA 95367; (209) 869-6633.

EMERGENCY MEDICINE. We are an established 35-physician partnership in northern California and we are seeking BE/BC Emergency Physicians to join us. All of our facilities have moderate volumes, many serve as EMS base-stations. Salary and benefits are competitive; malpractice is paid. If interested in a career in Emergency Medicine with us, please contact Sacramento Emergency Medical Group, 4325 Auburn Blvd, Ste 100, Sacramento, CA 95841; (916) 486-4414.

CARDIOLOGIST. BC/BE sought to join a large hospital-based multispecialty clinic. University affiliated residency program. Excellent fringe benefits. Competitive salary. California license required. Send CV to Aung-Win Chiong, MD, The Permanente Medical Group, Inc, PO Box 254999, Sacramento, CA 95865-4999, or call (916) 973-5748. An Equal Opportunity Employer

OREGON. ER Physician needed for small-town hospital located in recreational paradise. Modern facility serves as seven county referral center. Quick access to world-class windsurfing, superb snow skiing, and many other outdoor activities. 90 minutes from Portland. 12-hour shifts; \$75,000 plus with excellent benefits. Contact John Jacobson, MD, 1151 May St, Hood River, OR 97031; (503) 386-1111 (collect).

(Continued on Page 257)



IN HYPERTENSION

tance in the elderly

- □ Effective blood pressure control
- □ Low incidence of fatigue,^{2,3} impotence^{2,3} and cold extremities^{2,4}

Contraindicated in bronchial asthma, overt cardiac failure, greater-than-first-degree heart block, cardiogenic shock, and severe bradycardia.

See next page for references and Brief Summary of Product Information, which includes a listing of reported adverse reactions.



Because it also vasodilates

References: 1. Holtzman JL, Finley D, Johnson B, et al: The effects of single-dose atenolol, labetalol, and propranolol on cardiac and vascular function. Clin Pharmacol Ther 1986;40:268-273. 2. Due DL, Giguere GC, Plachetka JR: Postmarketing comparison of labetalol and propranolol in hypertensive patients. Clin Ther 1986;8(6):624-631. 3. Burris JF, Goldstein J, Zager PG, et al: Comparative tolerability of labetalol versus propranolol, aenolol, pindolol, metoprolol, and nadolol. J Clin Hypertens 1986;3:1-9. 4. Erb RJ, Plachetka JR: Thermographic evaluation of the peripheral vascular effects of labetalol and propranolol. Curr Ther Res 1985;28(1):68-73.

TRANDATE® Tablets

BRIEF SHMMARY OF PRODUCT INFORMATION

The following is a brief summary only. Before prescribing, see complete prescribing information in TRANDATE® Tablets product labeling.

IMDICATIONS AMD USAGE: TRANDATE® Tablets are indicated in the management of hypertension.

TRANDATE Tablets may be used alone or in combination with other antihypertensive agents, especially

CONTRAINDICATIONS: TRANDATE® Tablets are contraindicated in bronchial asthma, overt cardiac failure, CONTRAINDICATIONS: TRANDATE® Tablets are contraindicated in bronchial asthma, overt cardiac failure, greater-than-first-degree heart block, cardiogenic shock, and severe bradycardia (see WARNINGS). WARNINGS: Cardiac Failure: Sympathetic stimulation is a vital component supporting circulatory function in congestive heart failure. Beta-blockade carries a potential hazard of further depressing myocardial contractifity and precipitating more severe failure. Although beta-blockers should be avoided in vert congestive heart failure, if necessary labetalol HCl can be used with caution in patients with a history of heart failure who are well compensated. Congestive heart failure has been observed in patients receiving labetalol HCl. Labetalol HCl does not abolish the inotropic action of digitalis on heart muscle. In Patients Without a History of Cardiac Failure: In patients with latent cardiac insufficiency, continued depression of the myocardium with beta-blocking agents over a period of time can, in some cases, lead to cardiac failure. At the first sign or symptom of impending cardiac failure, patients should be fully digitalized and/or be given a diuretic, and the response should be observed closely. If cardiac failure continues despite adequate digitalization and diuretic. TRANDATE® theraoy should be withdrawn (gradually. if

despite adequate digitalization and diuretic, TRANDATE® therapy should be withdrawn (gradually, if

possible).

Exacerbation of Ischemic Heart Disease Following Abrupt Withdrawal: Angina pectoris has not been reported upon labetalol HCl discontinuation. However, hypersensitivity to catecholamines has been party to the property of the party of the par observed in patients withdrawn from beta-blocker therapy; exacerbation of angina and, in some cases, myocardial infarction have occurred after abrupt discontinuation of such therapy. When discontinuing myocardial infarction nave occurred arter abrupt discontinuation of such inerapy, miler discontinuing chronically administered TRANDATE, particularly in patients with ischemic heart disease, the dosage should be gradually reduced over a period of one to two weeks and the patient should be carefully monitored. If angina markedly worsens or acute coronary insufficiency develops, TRANDATE administration should be reinstituted promptly, at least temporarily, and other measures appropriate for the management of unstable angina should be taken. Patients should be warned against interruption or discontinuation of therapy without the physician's advice. Because coronary artery disease is common and may be unrecog-nized, it may be prudent not to discontinue TRANDATE therapy abruptly even in patients treated only for

Internation in many between the process of the process of the particular of the part beta-agonists is minimized.

Detail and interest in interest in the properties of the propertie

HCI to patients with pheochromocytoma.

Biabetes Mellitus and Hypoglycemia: Beta-adrenergic blockade may prevent the appearance of premonitory signs and symptoms (eg, tachycardia) of acute hypoglycemia. This is especially important with labile diabetics. Beta-blockade also reduces the release of insulin in response to hyperglycemia; it may there-

The description of the dose of antidiabetic drugs.

Major Surgery: The necessity or desirability of withdrawing beta-blocking therapy prior to major surgery is controversial. Protracted severe hypotension and difficulty in restarting or maintaining a heartbeat have been reported with beta-blockers. The effect of labetalol HCl's alpha-adrenergic activity has not

been evaluated in this setting.

A synergism between labetalol HCl and halothane anesthesia has been shown (see **Drug Interactions** under **PRECAUTIONS**).

PRECAUTIONS: General: Impaired Hepatic Function: TRANDATE® Tablets should be used with caution in patients with impaired hepatic function since metabolism of the drug may be diminished. Aundice or Hepatic Dysfunction: On rare occasions, labetalol HCl has been associated with jaundice (both hepatic and cholestatic). It is therefore recommended that treatment with labetalol HCl be stopped

(both hepatic and cholestatic). It is therefore recommended that treatment with labetalol HCl be stopped immediately should a patient develop jaundice or laboratory evidence of liver injury. Both have been shown to be reversible on stopping therapy.

Information for Patients: As with all drugs with beta-blocking activity, certain advice to patients being treated with labetalol HCl is warranted. This information is intended to aid in the safe and effective use of this medication. It is not a disclosure of all possible adverse or intended effects. While no incidence of the abrupt withdrawal phenomenon (exacerbation of angina pectoris) has been reported with labetalol HCl, dosing with TRANDATE Tablets should not be interrupted or discontinued without a physical sadvice. Patients being treated with TRANDATE Tablets should consult a physician at any sign of impending cardiac failure. Also, transient scalp tingling may occur, usually when treatment with TRANDATE Tablets is initiated (see ADVERSE REACTIONS).

Laboratory Tests: As with any new drug given over prolonged periods, laboratory parameters should be observed over regular intervals. In patients with concomitant illnesses, such as impaired renal function, appropriate tests should be done to monitor these conditions.

appropriate tests should be under the infinite trees conditions. Drug Interactions: In one survey, 2.3% of patients taking labetalol HCl in combination with tricyclic antidepressants experienced tremor as compared to 0.7% reported to occur with labetalol HCl alone. The contribution of each of the treatments to this adverse reaction is unknown, but the possibility of a drug

Drugs possessing beta-blocking properties can blunt the bronchodilator effect of beta-receptor agonist

Drugs possessing beta-blocking properties can blunt the bronchodilator effect of beta-receptor agonist drugs in patients with bronchospam; therefore, doses greater than the normal antiasthmatic dose of beta-agonist bronchodilator drugs may be required.

Cimetidine has been shown to increase the bioavailability of labetalol HCI. Since this could be explained either by enhanced absorption or by an alteration of hepatic metabolism of labetalol HCI, special care should be used in establishing the dose required for blood pressure control in such patients. Synergism has been shown between halothane anesthesia and intravenously administered labetalol HCI. During controlled hypotensive anesthesia using labetalol HCI in association with halothane, high concentrations (3% or above) of halothane should not be used because the degree of hypotension will be increased and because of the possibility of a large reduction in cardiac output and an increase in central venous pressure. The anesthesiologist should be informed when a patient is receiving labetalol HCI. Labetalol HCI blunts the reflex tachycardia produced by nitroglycerin without preventing its hypotensive effect. If labetalol HCI is used with nitroglycerin in patients with angina pectoris, additional antihyper-tensive effects may occur.

Drug/Laboratory Test Interactions: The presence of a metabolite of labetalol in the urine may result in falsely increased levels of urinary catecholamines when measured by a nonspecific trihydroxyindole (THI) reaction. In screening patients suspected of having a pheochromocytoma and being treated with labetalol HCI, specific radioenzymatic or high performance liquid chromatography assay techniques should be used to determine levels of catecholamines or their metabolites.

Carcinogenesis, Mutagenesis, Impairment of Fertility: Long-term oral dosing studies with labetalol HCl for 18 months in mice and for two years in rats showed no evidence of carcinogenesis. Studies with labetatol HCI using dominant lethal assays in rats and mice and exposing microorganisms according to modified Ames tests showed no evidence of mutagenesis.

Pregnancy: Reratogenic Effects: Pregnancy Category C: Teratogenic studies have been performed with

TRANDATE® Tablets (labetalol hydrochloride)

labetalol in rats and rabbits at oral doses up to approximately six and four times the maximum recommended human dose (MRHD), respectively. No reproducible evidence of fetal mafformations was observed. Increased fetal resorptions were seen in both species at doses approximating the MRHD. There are no adequate and well-controlled studies in pregnant women. Labetalol should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.

Nonteratogenic Effects: Infants of mothers who were treated with labetalol HCl during pregnancy did

not appear to be adversely affected by the drug. Oral administration of labetalol to rats during late gestation through weaning at doses of two to four times the MRHD caused a decrease in neonatal survival. Labor and Delivery: Labetalol HCl given to pregnant women with hypertension did not appear to affect the usual course of labor and delivery.

the usual course of labor and delivery.

Mursing Mothers: Small amounts of labetalol (approximately 0.004% of the maternal dose) are excreted in human milk. Caution should be exercised when TRANDATE Tablets are administered to a nursing woman.

Pediatric Use: Safety and effectiveness in children have not been established.

ADVERSE REACTIONS: Most adverse effects are mild, transient, and occur early in the course of treatment. In controlled clinical trials of three to four months' duration, discontinuation of TRANDATE® Tablets due to one or more adverse effects was required in 7% of all patients. In these same trials, beta-blocker control agents led to discontinuation in 8% to 10% of patients, and a centrally acting alpha-agonist in 30% of patients.

patients.

The following adverse reactions were derived from multi-center, controlled clinical trials over treatment periods of three and four months. The rates, which ranged from less than 1% to 5% except as otherwise noted, are based on adverse reactions considered probably drug-related by the investigator. If all reports are considered, the rates are somewhat higher (eg. dizziness, 20%; nausea, 14%; fatigue, 11%). Body as a Whole: Fatigue, asthenia, and headache.
Gastrointestinal: Nausea (6%), vomiting, dyspepsia, diarrhea, and taste distortion.
Central and Peripheral Mervous Systems: Dizziness (11%), paresthesia, and drowsiness.
Autonomic Nervous System: Nasal stuffiness, ejaculation failure, impotence, and increased

Cardiovascular: Edema and postural hypotension.

Respiratory: Dyspnea.

Special Senses: Vision abnormality and vertigo.
The adverse effects were reported spontaneously and are representative of the incidence of adverse effects that may be observed in a properly selected hypertensive patient population, ie, a group excluding patients with bronchospastic disease, overt congestive heart failure, or other contraindications to beta-

Clinical trials also included studies utilizing daily doses up to 2,400 mg in more severely hypertensive patients. The US therapeutic trials data base for adverse reactions that are clearly or possibly dose-related shows that the following side effects increased with increasing dose: dizziness, fatigue, nausea, vomiting, dyspepsia, paresthesia, nasal stuffiness, ejaculation failure, impotence, and edema.

In addition, a number of other less common adverse events have been reported in clinical trials or the

Cardiovascular: Postural hypotension, including rarely, syncope.

Central and Peripheral Nerrous Systems: Paresthesias, most frequently described as scalp tingling. In most cases, it was mild, transient, and usually occurred at the beginning of treatment.

Collagen Disorders: Systemic lupus erythematosus; positive antinuclear factor (ANF).

Immunological System: Antimitochondrial antibodies.
Liver and Biliary System: Cholestasis with or without jaundice.

Musculoskeletal System: Muscle cramps; toxic myopathy.

Respiratory System: Bronchospasm.

NSim and Appendages. Rashes of various types, such as generalized maculopapular, lichenoid, urticarial, bullous lichen planus, psoriaform, and facial erythema; Peyronie's disease; reversible alopecia. Urinary System: Difficulty in micturition, including acute urinary bladder retention. Following approval for marketing in the United Kingdom, a monitored release survey involving approximately 6,800 patients was conducted for further safety and efficacy evaluation of this product. Results of this survey indicate that the type, severity, and incidence of adverse effects were comparable to those

Potential Adverse Effects: In addition, other adverse effects not listed above have been reported with

other beta-adrenergic blocking agents.

Central Nervous System: Reversible mental depression progressing to catatonia, an acute reversible syndrome characterized by disorientation for time and place, short-term memory loss, emotional lability, slightly clouded sensorium, and decreased performance or neuropsychometrics. Cardiovascular: Intensification of AV block (see CONTRAINDICATIONS).

Allergic: Fever combined with aching and sore throat; laryngospasm; respiratory distress. Hematologic: Agranulocytosis; thrombocytopenic or nonthrombocytopenic purpura. Gastrointestinal: Mesenteric artery thrombosis; ischemic colitis.

The oculomucocutaneous syndrome associated with the beta-blocker practolol has not been reported with labetalol HCI.

Clinical Laboratory Tests: There have been reversible increases of serum transaminases in 4% of patients treated with labetalol HCl and tested, and more rarely, reversible increases in blood urea. OVERDOSAGE: Information concerning possible overdosage and its treatment appears in the full prescrib-

ing information.

Dosage and administration: Dosage Must be individualized. The recommended *initial* dosage is DOSAGE AMN ADMINISTRATION: DOSAGE MUST BE INDIVIDUALIZED. The recommended initial dosage is 100 mg twice daily whether used alone or added to a diuretic regimen. After two or three days, using standing blood pressure as an indicator, dosage may be titrated in increments of 100 mg bid every two or three days. The usual maintenance dosage of labetalol HCI is between 200 and 400 mg twice daily. Before use, see complete prescribing information for dosage details.

HOW SUPPLIED: TRANDATE Tablets and to 100 mg, light orange, round, scored, film-coated tablets engraved on one side with "TRANDATE 100 GLAXO" bottles of 100 (NDC 0173-0346-43) and 500 (NDC 0173-0346-44) and unit dose packs of 100 tablets (NDC 0173-0346-47).

TRANDATE Tablets, 200 mg, white, round, scored, film-coated tablets engraved on one side with "TRANDATE 200 GLAXO" bottles of 100 (NDC 0173-0347-43) and 500 (NDC 0173-0347-44) and unit dose packs of 100 tablets (NDC 0173-0347-47).

TRANDATE Tablets, 300 mg, peach, round, scored, film-coated tablets engraved on one side with "TRANDATE 300 GLAXO" bottles of 100 (NDC 0173-0348-43) and 500 (NDC 0173-0348-44) and unit dose packs of 100 tablets (NDC 0173-0348-47).

TRANDATE Tablets, should be stored between 2° and 30°C (36° and 86°F). TRANDATE Tablets in the unit

packs on 100 capiets (NDC 01/3-0346-47).

TRANDATE Tablets should be stored between 2° and 30°C (36° and 86°F). TRANDATE Tablets in the unit dose boxes should be protected from excessive moisture.

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September 1986



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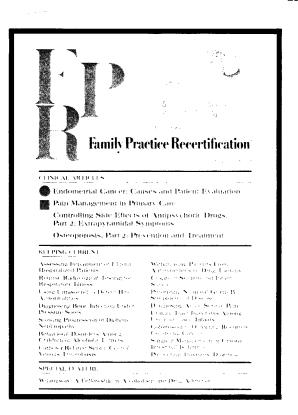
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PRACTICAL - CLINICAL - EDUCATIONAL - CURRENT

(Continued from Page 252)

PHYSICIANS WANTED

FAMILY PRACTICE PHYSICIANS. Discover the magic of the southwest. Experience New Mexicol Outstanding career opportunities for BC/BE, Family Practitioners. Obstetrics preferred, malpractice paid. Contact your Physician Consultant at New Mexico Health Resources, PO Box 27650, Albuquerque, NM 87125; (505) 242-0633. No fee.

INTERNIST. Immediate opening for a BC/BE General Internist to join a 35-member multispecialty group located in San Luis Obispo on the central coast. Benefits include immediate shareholder status, retirement program, all practice costs paid, starting guaranteed salary plus strong incentive plan. Send CV to Recruitment, San Luis Medical Clinic, Ltd., 1235 Osos St, San Luis Obispo, CA 93401-3619.

OREGON. Family Practice Physician partner needed for immediate care center and private practice in recreational Oregon paradise. No start-up costs, thriving practice, and generous free time to spend skiing or hiking the mountains or on the river, enjoying some of the best windsurfing waters in the world. Located in Hood River which offers the peace and quiet of a small community, yet is only an hour away from the culture and activity of Oregon's largest metropolitan area. Contact John Jacobson, MD, Medical Director, Care Corner, 1151 May St, Hood River, OR 97031; (503) 386-1111 (collect).

CALIFORNIA, SONORA. Staff Physician position available in 11-12,000 visit ER in quaint, historic, growing gold country community with fantastic recreational opportunities, one hour from Yosemite. Excellent opportunity in an academic and democratic group. Send CV to Art B. Wong, MD, FACEP, EPMG, 1 Maritime Plaza, Ste 710, San Francisco, CA 94111.

PHYSICIANS WANTED

SOUTHERN CALIFORNIA

Enjoy professional challenge and growth with a successful and expanding HMO in southern California. CIGNA Healthplans of California is seeking Specialists and Primary Care physicians committed to concepts of prevention and health maintenance to join our facilities in Los Angeles and Orange Counties. We offer an excellent compensation and benefits package including profit sharing. For consideration, please forward CV to:

Director/Physician Recruitment CIGNA Healthplans of California 505 N. Brand Blvd, Suite 400-49 Glendale, CA 91203

REDWOOD COUNTRY-PRIMARY CARE. Small multispecialty group seeks a Primary Care Physician, BC in Family Practice or Internal Medicine for a growing practice. Arcata is a coastal community in redwood country, offering a unique combination of rural lifestyle in a university town. Competitive salary and benefits package leading to partnership. Send CV to Arcata Family Medical Group, 4555 Valley West Blvd, Arcata, CA 95521.

THREE FULL-TIME GENERAL INTERNISTS are being recruited for positions in the Division of General Medicine, Department of Internal Medicine, University of New Mexico School of Medicine. Primary responsibilities include inpatient and outpatient care and teaching. Applicants with subspecialty training will be considered. Applicants must be BC/BE. Faculty appointment based on academic credentials and experience. Please send inquiries and résumé/CV to Pat Cleve, Department of Medicine, University of New Mexico, Albuquerque, NM 87131.

INTERNIST BC/BE to join Internist/Cardiologist in central California near Fresno. Reply to Number 85, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602.

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FAMILY PRACTICE PHYSICIAN. Immediate openings in multispecialty Community Health Center. Urban/rural sites available in Kern County, California. Very competitive salary and benefit package available. Modern facilities with comprehensive support system of lab, x-ray, pharmacy, etc. We believe we can offer a physician a supportive professional environment in which to practice while also offering an exceptional financial package. We also have the flexibility with five clinic sites available to provide a wide range of life-style and practice opportunities. For more information call Jack Hicks, Assistant Director, Clinica Sierra Vista, PO Box 457, Lamont, CA 93241; (805) 845-3731.

CARDIOLOGIST. Columbus Hospital in Great Falls, Montana, has an immediate need for a BE/BC, invasive/non-invasive Cardiologist. At the foot of the eastern slopes of the Rockies, Great Falls is best known for its accessibility to superb recreational activities—skiing, fishing, hunting, camping, hiking, back-packing, etc. With excellent school systems and a very moderate cost of living, Great Falls has an excellent overall quality of life. Opportunity offers a guaranteed income plus incentives. Please send CV to Charles Matenaer, PRM, Inc, 15975 W. National Ave, New Berlin, WI 53151, or call (414) 784-2777.

ORTHOPEDIST. The west coast's leading Occupational/Family Practice medical provider has FT/PT opportunities for Orthopedic Specialists in California and Washington (Seattle/Tacoma). Attractive package includes: guaranteed salary, incentive bonus and benefits. Current license. Contact Personnel Director, ReadiCare, Inc, 446 Oakmead Parkway, Sunnyvale, CA 94086; (800) 237-3234. Join our dynamic team of professionals. Practice and live in an incomparable environment.

(Continued on Page 263)

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California Medical Association's
117th Annual Session

THE WESTERN SCIENTIFIC ASSEMBLY

March 4-9, 1988
Bally's Hotel and Reno/Sparks Convention Center





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The



Telfer B. Reynolds, MD/Golden Apple Award

Telfer B. Reynolds, MD, Clayton G. Loosli Professor of Medicine at the University of Southern California, has been selected by the Committee on Scientific Assemblies to receive CMA's 1988 Golden Apple Award. This award spotlights exceptional physician speakers who have made a lifelong commitment to teaching and become distinguished by their charismatic, scientific and educational talents.



California Medical Association's 1988 Western Scientific Assembly will be held March 4-6 at Reno-Sparks Convention Center. Persons attending scientific courses or special conferences are asked to register on site in the North Hall Concourse of the convention center. For more information on the following programs please contact CMA at (415) 541-0900.

RIDAY, MARCH 4

D. D. et a. C. et dia a Life Common and W/a ulasha and
Basic Cardiac Life Support Workshop
☐ AIDS—What the Practicing Physican Should
Know in 1988
Techniques & Problems in Handling Hazardous
Material Disasters
☐ Sleep Disorders
Symposium on Developmental Disabilities
Pulmonary Medicine Update
Neurosurgery in the Aging Population
Golden Apple Award: Telfer B. Reynolds, MD

SATURDAY, MARCH 5

Advanced Cardiac Life Support:
Certification Course
☐ Cardiovascular Risk-Assessment and
Intervention in Primary Care
☐ What's Allergy, What's Not: Common Health
Problems in the Workplace
☐ Implantable Electromagnetic Hearing Device
Breast Cancer: Current Diagnosis and Treatment
Low Back Pain: Office Evaluation
and Management
Hair Symposium — Answer Your
Patients' Questions
Update on Gynecologic Cancer Screening
☐ It's A Wilderness Out There! Medical
Emergencies of the Great Outdoors
Opthalmology in the Year 2001
Colorectal Carcinoma: Update 1988
☐ Family Crisis of the 1980's: Alcohol and
Drug Abuse
☐ Eye Care in the Elderly

☐ The Impaired Doctor

Western Scientific Assembly



Dame Sheila Sherlock, DBE, MD, FACP Roberta F. Fenlon Memorial

One of the world's foremost authorities on liver disease, Dame Sheila Sherlock is the renowned author of *Diseases of the Liver and Biliary System* and numerous scientific papers on hepatic pathology, function and circulation.



Robert S. Eliot, MD, FACC Keynote Speaker of the Western Scientific Assembly

Author of the popular book on stress, *Is It Worth Dying For?*, Doctor Robert Eliot is nationally known for his research on the effects of lifestyle, behavior and stress on health, particularly heart disease.

SATURDAY, MARCH 5

- Skin Cancer: Everyone's Problem
- ☐ The Neuropsychiatrically Impaired Physician
- ☐ Marketing Roundtable 1988
- Roberta F. Fenlon Memorial Lecture:

Dame Sheila Sherlock

SUNDAY, MARCH 6

- Advanced Cardiac Life Support: Certification/Recertification Course
- What the Non-Urologist Should Know About
- Urology in '88
 ☐ Recognition of Premalignant and Precursor
- Lesions of the Skin and Genitalia
- ☐ Suture Course
- ☐ Adolescent Psychological Problems
- ☐ Seminars in Orthopaedic Management
- ☐ What Every Computer Illiterate Should Know
- ☐ Consulting with the Anesthesiologist
- ☐ A Critical Look at the Washington Scene
- ☐ Public Health and Policy: AIDS in California 1988
- ☐ Physician and the Law 1988
- ☐ Keynote Speaker: Robert S. Eliot, MD, FACC



The Western Scientific Assembly

CONCURRENT EVENTS

HOUSE OF DELEGATES

The CMA House of Delegates meets Saturday, March 5 through Wednesday, March 9 at Bally's Hotel. Sit in on reference committee hearings and House sessions to see firsthand how CMA policies are shaped. Highlights of this year's House will be the farewell address of out-going CMA President Dr. Frederick S. Armstrong, a San Jose internist, and the inaugural address of CMA's President-Elect, Dr. Laurens P. White, a San Francisco internist specializing in oncology. Registration will be available March 3-9 at Bally's Hotel in the Grand Salon.

HOSPITAL MEDICAL STAFF SECTION ANNUAL ASSEMBLY

The CMA-HMSS Annual Assembly will be Friday, March 5 at Bally's Hotel. Included will be their keynote speaker, Kenneth Nelson, MD, "The View from Here—the PRO Sanction Process."

SPECIAL EVENTS

A variety of special tours will be offered this year. Regis-
tration takes place at the Reno-Sparks Conven-
tion Center.

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- ☐ Virginia City Tour
- ☐ Lake Tahoe Cruise
- Western Hoe-Down and Harrah's Classic Auto Tour
- ☐ Pre and Post Convention Ski Packages:
 - Ski Lake Tahoe

EXHIBITS

The exhibit hall at Reno-Sparks Convention Center offers something for everyone: pharmaceuticals, educational displays, computer and office systems, medical testing systems, insurance, investment opportunities, health and home helps. Prizes!

GENERAL INFORMATION

LOCATION

The CMA 1988 Western Scientific Assembly will be held at the Reno-Sparks Convention Center, 4590 S. Virginia Street, Reno, Nevada, 89502. The House of Delegates will be headquartered in the Bally's Reno Hotel, 2500 E. Second Street, Reno, Nevada, 89595. Call (702) 789-2033 for hotel reservation information.

FEES

There is no charge to CMA members and their non-physician family members for general registration, nor to registered nurses, residents, interns or medical students. A general registration fee of \$150 is charged to non-member physicians.

REGISTRATION

Advance registration closes February 12, 1988. Attendees may register on site at the Reno/Sparks Convention Center or Bally's Reno Hotel. Registration will be open:

Thursday, March 3

1-5 pm Bally's Hotel, Grand Salon

Friday, March 4

7:30 am-5 pm Bally's Hotel, Grand Salon 7 am-5 pm Convention Center

Saturday, March 5

7 am-5 pm Bally's Hotel, Grand Salon 7 am-5 pm Convention Center

Sunday, March 6

7:30 am-5 pm Bally's Hotel, Grand Salon

7 am-5 pm Convention Center

SHUTTLE BUS SERVICE

Shuttle service from Bally's Hotel to Reno-Sparks Convention Center (located 3 miles away) will be continuous from 6:30 am to 7:30 pm.

ANY QUESTIONS?

Call CMA, (415) 541-0900

(Continued from Page 257)

PHYSICIANS WANTED

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Medical Director Laguna Honda Hospital 375 Lagunda Honda Blvd San Francisco, CA 94116

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ARIZONA INTERNAL MEDICINE PHYSICIAN to associate with two man Internal Medicine group in Tucson. New office building with lab and x-ray. Well established practice. Sub-specialties welcome, BC/BE. Send CV to A. Oaks, 5265 E. Knight Dr, Tucson, AZ 85712.

FAMILY PHYSICIAN needed for private practice opportunity in southeast New Mexico community. Excellent potential for thriving practice. Contact NM Health Resources, PO Box 27650, Albuquerque, NM 87125; (505) 242-0633.

PHYSICIANS WANTED. A General Surgeon and an Oncologist to join 16-physician multispecialty group with attached 40-bed hospital located in southern Idaho. Contact Business Administrator, Box 1233, Twin Falls, ID 83301.

OUTSTANDING OPPORTUNITIES. Oncologist with Internal Medicine strength and General Internist, BE/BC, private practice or affiliation available, San Francisco north bay area community. Send résumé to Number 84, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602.

NORTHERN CALIFORNIA rural community clinic seeks BC/BE Family Physician. Heart of the redwoods. Full range of practice, community hospital nearby. Contact Allan Katz, (707) 923-2783.

INVASIVE/NONINVASIVE CARDIOLOGIST. Fulltime Cardiologist needed. Salaried position with excellent benefits. Send CV to Lancaster Cardiology Medical Group, Inc, 43847 N. Heaton Ave, Lancaster, CA 93534.

CALIFORNIA, SAN FRANCISCO BAY AREA. BC/BE OB/GYN. Multiracial community clinic needs part-time OB/GYN for ambulatory and surgical services, ten to twenty deliveries monthly, many high risk. Close to downtown San Francisco and Oakland. Excellent opportunity for new graduate starting practice or early retiree. \$80,000 annually, plus paid malpractice. Contact Alice Godfrey, MD, 101 Broadway, Richmond, CA 94804; (415) 233-3994.

LOCUM TENENS opportunities: temporary or permanent. Guaranteed income/paid malpractice. Contact Diane Smithline, Current Health Concepts, Inc, Locum Tenens Division, 100 Woodward Ave, Sausalito, CA 94965; (415) 331-7422. Physician owned and operated.

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FELLOWSHIP IN GERIATRIC MEDICINE. A unique fellowship in Geriatric Medicine is being offered beginning July 1, 1988, by a Fortune 100 company and a major university medical school in California. The program is two years in duration and provides the gamut of geriatric health care experience and qualifies one for the certificate examination in Geriatric Medicine. The fellowship stipend is attractive as are the working conditions in this unique experience. For further information please contact Number 86, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602.

EXCELLENT TEXAS OPPORTUNITIES in Cardiology, ENT, Family Practice, General Surgery, Internal Medicine, OB/GYN, Oncology, Orthopedic Surgery, Vascular Surgery, Urology, Industrial Medicine. Excellent quality of life, first year guarantee, etc. Reply with CV or call Armando L. Frezza, Medical Support Services, 8806 Balcones Club Dr, Austin, TX 78750; (512) 331-4164.

PACIFIC NORTHWEST. BE/BC Pediatrician wanted to join two Pediatricians in busy migrant health clinic. Family oriented community. Rural lifestyle. Excellent recreation. Negotiable salary, includes malpractice. Contact Ann Garza, YVFWC, PO Box 190, Toppenish, WA 98948; (509) 865-5600.

Western States OPENINGS

Many multispecialty groups and hospitals have asked us to recruit for over 300 positions of various specialties. Send CV to:

Western States Physician Services 407 S. Clovis Ave, Ste 108, Fresno, CA 93727 Or call (209) 252-3000

SITUATION WANTED

PLASTIC SURGEON. Completed residency in Detroit and suburbs. Form a win-win situation with you having time off covered, and increasing your net income, and with me locating in a favorable spot. Please reply to Number 82, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602.

MD-MPH (Health Services Admin) seeks nonclinical management position in provider and/or non-provider setting in Seattle area. Extensive experience at all levels of UR and QA. Very experienced with federal and local accreditation and regulatory agencies. Also have physician management experience. Superb communication and presentation skills. CV, references, and interview in confidence on request. Reply to Number 83, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602.

RADIOLOGIST. BC for locum/part-time or full-time. (209) 931-4357: evenings, weekends, some days.

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Western Physicians Registry 710 Van Ness Ave San Francisco, CA 94102

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PRACTICES AVAILABLE

BEAUTIFUL PUGET SOUND FAMILY PRACTICE available. Complete office, lab and x-ray equipment. Practice gross over 225K. Price \$35,000. Reply to Number 74, Western Journal of Medicine, PO Box 7602, San Francisco, CA 94120-7602.

SOUTH SACRAMENTO AREA FAMILY PRACTICE. Collections of \$111K in 1986. 95% insured patients. Full price: \$50K. Seller financing available. Call Western Practice Sales (916) 673-1302.

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MEDICAL OFFICE SPACE AVAILABLE. Next to Desert Samaritan Hospital, Mesa, Arizona. Partly furnished. Vacant two and one-half days a week or can be shared with doctor additional two and one-half days. Phone (602) 254-5835.

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(Continued on Page 264)

(Continued from Page 263)

MEDICAL PRACTICE OPPORTUNITIES

CALIFORNIA/NATIONWIDE. IM, Gastro, Pulm, Cardio, Ortho, Surg, FP, Nutrition, Derm, Psych, Neurol, OB/GYN, Oph, ENT, Urol, plus others. BRADSHAW ASSOCIATES, Practice Sales/Recruitment/Valuations, 21 Altamount Dr, Orinda, CA 94563; (415) 376-0762.

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New $24' \times 36'$ three car garage with wood stove and $24' \times 40'$ outdoor garage and shop, both with full foundations and cement floors, $16' \times 40'$ horse barn-tack room-hay shed.

Full redwood decks on three sides of house, double post and three rail fencing on 16' centers around the 10 acres.

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MARGARET ROSS

PHYSICIAN PLACEMENT COORDINATOR

CALIFORNIA MEDICAL ASSOCIATION

221 Main St, PO Box 7690, San Francisco, CA 94120-7690 (415) 541-0900

References: 1. BAC-DATA Medical Information Systems, Inc., Volume I, 1986. 2. Schwartz RH, et al. Rev Infect Dis 4:514-516, 1982. 3. Data on file. Hoffmann-La Roche Inc., Nutley, NJ. 4. Drug Topics Red Book

BACTRIM'" (trimethoprim and sulfamethoxazole/Roche)

Before prescribing, please consult complete product information, a summary of which follows:
CONTRAINDICATIONS: Hypersensitivity to trimethoprim or sulfonamides, documented megaloblastic

anemia due to folate deficiency, pregnancy at term and during the nursing period; infants less than two months of age.

WARNINGS: FATALITIES ASSOCIATED WITH THE ADMINISTRATION OF SULFONAMIDES, ALTHOUGH RARE, HAVE OCCURRED DUE TO SEVERE REACTIONS, INCLUDING STEVENS-JOHNSON SYNDROME, TOXIC EPIDERIMAL NECROLYSIS, FULMINANT HEPATIC NECROSIS, AGRANULOCYTOSIS, APLASTIC AMEMIA AND OTHER BLOOD DYSCRASSIS.

BACTRIM SHOULD BE DISCONTINUED AT THE FIRST APPEARANCE OF SKIN RASH OR ANY SIGNED ADVERSE BEACTION. Cliniques signs, such as resh, some threat fewer called purpour agriculture, may be

ADVERSE REACTION. Clinical signs, such as rash, sore throat, fever, pallor, purpura or jaundice, may be early indications of serious reactions. In rare instances a skin rash may be followed by more severe reactions, such as Stevens-Johnson syndrome, toxic epidermal necrolysis, hepatic necrosis or serious blood

disorder. Perform complete blood counts frequently.

BACTRIM SHOULD NOT BE USED IN THE TREATMENT OF STREPTOCOCCAL PHARYNGITIS. Clinical studies show that patients with group A 6-hemolytic streptococcal tonsilopharyngitis have a greater incidence of bacteriologic failure when treated with Bactrim than with penicillin.

PRECAUTIONS: General: Give with caution to patients with impaired renal or hepatic function, possible folate deficiency (e.g., elderly, chronic alcoholics, patients on anticonvulsants, with malabsorption syndrome, or in mainutrition states) and severe allergies or bronchial asthma. In glucose-6-phosphate dehy-

drome, or in malnutrition states) and severe allergies or bronchial asthma. In glucose-6-phosphate dehydrogenase delicient individuals, hemolysis may occur. I requently dose-related. Use in the Elderly: May be increased risk of severe adverse reactions in elderly, particularly with complicating conditions, e.g., impaired kidney and/or liver function, concomitant use of other drugs. Severe skin reactions, generalized bone marrow suppression (see WARNINGS and ADVERSE REACTIONS) or a specific decrease in platelets (with or without purpura) are most frequently reported severe adverse reactions in elderly. In those concurrently receiving certain diuretics, primarily thiazides, increased incidence of throm-bocytopenia with purpura reported. Make appropriate dosage adjustments for patients with impaired kidney function (see DOSAGE AND ADMINISTRATION).

function (see DOSAGE AND ADMINISTRATION).

Use in the Treatment of Pneumocystis Carinii Pneumonitis in Patients with Acquired Immunodeficiency Syndrome (AIDS). Because of unique immune dysfunction, AIDS patients may not tolerate or respond to Bactrim in same manner as non-AIDS patients. Incidence of side effects, particularly rash, fever, leukopenia, with Bactrim in AIDS patients treated for Pneumocystis carinii pneumonitis reported to be greatly increased compared with incidence normally associated with Bactrim in non-AIDS patients.

Information for Patients: Instruct patients to maintain adequate fluid intake to prevent crystalluria and stone

Individual of Paleins. Institute patients to maintain adequate into intake to prevent crystainina and stone Laboratory Fests: Perform complete blood counts frequently; if a significant reduction in the count of any formed blood element is noted, discontinue Bactrim. Perform urinalyses with careful microscopic examination and renal function tests during therapy, particularly for patients with impaired renal function.

Drug Interactions: In elderly patients concurrently receiving certain diuretics, primarily thiazides, an increased incidence of thrombocytopenia with purpura has been reported. Bactrim may prolong the prothrombin time in patients who are receiving the anticoagulant warfarin. Keep this in mind when Bactrim is given to patients already on anticoagulant therapy and reassess coagulation time. Bactrim may inhibit the hepatic metabolism of phenytoin. Given at a common clinical dosage, it increased the phenytoin half-life by 39% and decreased the phenytoin math-life by 39% and decreased the phenytoin half-life by Drug/Laboratory fest Interactions: Bactrim as a protein binding sites, thus increasing free methotrexate concentrations.
Drug/Laboratory fest Interactions: Bactrim, specifically the trimethoprim component, can interfere with a serum methotrexate assay as determined by the competitive binding protein technique (CBPA) when a sacterial dihydrofolate reductase is used as the binding protein. No interference occurs if methotrexate is measured by a radioimmunoassay (RIA). The presence of trimethoprim and sulfamethoxazole may also interfere with the Jaffé alkaline picrate reaction assay for creatinine, resulting in overestimations of about 10% in the range of normal values.

10% in the range of normal values.

Carcinogenesis, Mutagenesis, Impairment of Fertility: Carcinogenesis: Long-term studies in animals to evaluate carcinogenic potential not conducted with Bactrim. Mutagenesis: Bacterial mutagenic studies not performed with sulfamethoxazole and trimethoprim in combination. Trimethoprim demonstrated to be performed with sulfamethoxacole and trimethoprim in combination. Trimethoprim demonstrated to be normutagenic in the Ames assay. No chromosomal damage observed in human leukocytes in vitro with sulfamethoxazole and trimethoprim alone or in combination; concentrations used exceeded blood levels of these compounds following therapy with Bactim. Observations of leukocytes obtained from patients treated with Bactim revealed no chromosomal abnormatities. Impairment of Fertility: No adverse effects on fertility or general reproductive performance observed in rats given oral dosages as high as 70 mg/kg/day trimethoprim plus 350 mg/kg/day sulfamethoxazole. Pregnancy: Teratogenic Effects: Pregnancy Category C. Trimethoprim and sulfamethoxazole may interfere with folic acid metabolism. use during pregnancy only if potential benefit justifies potential risk to letus. Nonteratogenic Effects: See CONTRAINDICATIONS section.

Nursing Mothers: See CONTRAINDICATIONS section.

Nursing Mothers: See CONTRAINDICATIONS section.

Pediatric Use: Not recommended for infants under two months (see INDICATIONS and CONTRAINDICA

TIONS SECTIONS). MOST common are gastrointestinal disturbances (nausea, vomiting, anorexia) and allergic skin reactions (such as rash and urticaria). FATALITIES ASSOCIATED WITH THE ADMINISTRATION OF SULFONAMIDES, ALTHOUGH RARE, HAVE OCCURRED DUE TO SEVERE REACTIONS, INCLUDING STEVENS-JOHNSON SYNDROME, TOXIC EPIDERMAIL NECROLYSIS, FULLMINANT HEPATIC MECROSIS, AGRAMULOCYTOSIS, APLASTIC ANEMIA AND OTHER BLOOD DYSCRASIAS (SEE WARNINGS SECTION). STEVENS-JOHNSON SYNDRÖME, TOXIĆ EPIDERMAL NECROLYSIS, FULMINANT HEPATIC MECROSIS, AGRAMULOCYTOSIS, APLASTIC ANEMIA AND OTHER BLOOD DYSCRASIAS (SEE WARNINGS SECTION). Hematologic: Agranulocytosis, aplastic anemia, thrombocytopenia, leukopenia, neutropenia, hemolytic anemia, megaloblastic anemia, hypoprothrombinemia, methemoglobinemia, eosinophiia. Allergic Réactions: Stevens-Johnson syndrome, toxic epidermal necrolysis, anaphylaxis, allergic myocarditis, erythema multiforme, exfoliative dermatitis, angioedema, drug fever, chills, Henoch-Schoeniein purpura, serum sickness-like syndrome, generalized allergic reactions, generalized sick in eruptions, photosensitivity, conjunctival and scleral injection, pruritus, urticaria and rash. Periarteritis nodosa and systemic lupus erythematosus have been reported. Gastrointestinal: Hepatitis (including cholestatic jaundice and hepatic necrosis), elevation of serum transaminase and bilirubin, pseudomembranous enterocolitis, pancreatitis, stomatitis, glossitis, nausea, emesis, abdominal pain, diarrhea, anorexia. Genitourinary: Renal failure, interstitial nephritis, BUM and serum creatinine elevation, toxic nephrosis with oliquira and anuria, crystal-luria. Neurologic: Aseptic meningitis, convulsions, peripheral neuritis, ataxia, vertigo, tinnitus, headache. Psychiatric: Hallucinations, depression, apathy, nervousness. Endocrine: Sulfonamides bear certain chemical similarities to some gottrogens, diuretics (acetazolamide and the thiazides) and oral hypoglycemic agents. cross-sensitivity may exist. Duresis and hypoglycemia have occurred rarely in patients receiving sulfonamides. Musculosketela: Arthraligia, myalgia. Miscellaneous: Weakness, fatigue, insomnia.

DOSAGE AND ADMINISTRATION: Not recommended for use in infants less than two months of age.
URINARY TRACT INFECTIONS AND SHIGELLOSIS IN ADULTS AND CHILDREN. AND ACUTE CTINTS MEDIA IN CHILDREN. AND

sulfamethoxazole per 24 hours in equal doses every 6 hours for 14 days. See complete product information for suggested children's dosage table.

HOW SUPPLIED: DS (double strength) Tables (160 mg trimethoprim and 800 mg sulfamethoxazole)—bottles of 100, 250 and 500; Tel-E-Dose* packages of 100; Prescription Paks of 20. Tables (80 mg trimethoprim and 400 mg sulfamethoxazole)—bottles of 100 and 500. Tel-E-Dose* packages of 100; Prescription Paks of 40. Pediatric Suspension (40 mg trimethoprim and 200 mg sulfamethoxazole per teasp.)—bottles of 100 mand 16 oz (1 pint). Suspension (40 mg trimethoprim and 200 mg sulfamethoxazole per teasp.)—bottles of 16 oz (1 pint). Suspension (40 mg trimethoprim and 200 mg sulfamethoxazole per teasp.)—bottles of 16 oz (1 pint).

STORE TABLETS AT 15°-30°C (59°-86°F) IN A DRY PLACE PROTECTED FROM LIGHT. STORE SUSPENSIONS AT 15°-30°C (59°-86°F) PROTECTED FROM LIGHT.



The power persists.

- Succeeds year after year—with little change in resistance
- Succeeds against organisms common to otitis media *in vitro*
- Succeeds with b.i.d. convenience and pleasant cherry flavor

From year to year, Bactrim destroys not only *H. influenzae* and *S. pneumoniae in vitro*, but also various ampicillin-resistant strains. ^{2.3} And *b.i.d.* Bactrim costs less than *t.i.d.* cefaclor or *t.i.d.* amoxicillin/clavulanate potassium. ⁴

In vitro results may not correlate with clinical results. Not indicated for prophylactic or prolonged use in otitis media and contraindicated in infants under two months of age.

Specify. Do Not Substitute.

Bactrim[®] Pediatric

(40 mg trimethoprim and 200 mg sulfamethoxazole per 5 ml)

It keeps its powers.